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Amendments to the Specification:

At page 13, lines 16-30, please replace paragraph with the following:

The Expect value is used as a convenient way to create a significance threshold for reporting results. The default value used for blasting is typically set to 0.0001. In BLAST 2.0, the Expect value is also used instead of the P value (probability) to report the significance of matches. For example, an E value of one assigned to a hit can be interpreted as meaning that in a database of the current size one might expect to see one match with a similar score simply by chance. An E value of zero means that one would not expect to see any matches with a similar score simply by chance. See, e.g., http://www.nebi.nlm.nih.gov/Education/BLASTinfo/. Occasionally, a string of X's or N's will result from a BLAST search. This is a result of automatic filtering of the query for low-complexity sequence that is performed to prevent artifactual hits. The filter substitutes any low-complexity sequence that it finds with the letter "N" in nucleotide sequence (e.g., "NNNNNNNNNNNNNNNN") or the letter "X" in protein sequences (e.g., "XXXXXXXXXXX"). Low-complexity regions can result in high scores that reflect compositional bias rather than significant position-by-position alignment. (Wootton and Federhen, Methods Enzymol 266:554-571, 1996).

At page 23, please replace Table 1J with the following:

| Table 1J. BLAST results for NOV1a | | | | | |
|--|--|----------------|------------------|------------------|--------|
| Gene Index/ Identifier | Protein/ Organism | Length (aa) | Identity (%) | Positives (%) | Expect |
| gi 13569928 ref NP_ 112217.1 NP 112217 | a disintegrin-like and metalloprotease with thrombospondin type 1 motif, 12 [Homo sapiens] | 1593 | 269/616 (43%) | 371/616 (59%) | e-135 |
| gi 11493589 gb AAG3 5563.1 AF163762_1 (AF163762) AAG35563 | zinc metalloendopeptida se [Homo sapiens] | 1077 | 253/624 (40%) | 359/624 (56%) | e-128 |

| | | , | | | , |
|----------------------|--------------------|--------------|-------------|---------|--------------|
| gi 10645199 ref NP | a disintegrin and | 997 | 247/571 | 344/571 | e-127 |
| 055087.1 | metalloprotease | | (43%) | (59%) | |
| NP 055087 | with | | | | 1 |
| | thrombospondin | | | | |
| | motifs-7 | | | | |
| İ | preproprotein; a | | | | |
| | disintegrin-like | [| | | |
| | and | | | | 1 |
| | metalloprotease | | | | |
| | (reprolysin type) | | | | |
| | with | | | | 1 |
| | thrombospondin | | | |] |
| | type 1 motif, 7 | i | | | · |
| | [Homo sapiens] | | | | |
| gi 15309931 ref XP | a disintegrin-like | 854 | 245/604 | 348/604 | e-124 |
| 054419.1 | and | | (40%) | (57%) | |
| XP 054419 | metalloprotease | | ,, | (2,0) | 1 |
| | domain with | ľ | | | |
| | thrombospondin | | | | |
| | type I repeats 10 | | | | |
| | [Homo sapiens] | | | | |
| gi 7656869 ref NP 0 | a disintegrin and | 860 | 244/630 | 345/630 | e-115 |
| 55088.1 NP 055088 | metalloprotease | | (38%) | (54%) | |
| 3333311 | with | | ``` | (310) | |
| | thrombospondin | | , | | |
| | motifs-6 | | | | |
| | preproprotein; a | | | | |
| | disintegrin-like | | | | |
| | and | | | | |
| | metalloprotease | | | | |
| | (reprolysin type) | | | | |
| | with | | | | |
| | thrombospondin | | | | |
| | type 1 motif, 6 | | | t. | |
| | [Homo sapiens] | | | | |
| | [110000 sabiens] | | l | | |

At page 24 please replace Table 1K with the following:

Table 1K. ClustalW Analysis of NOV1

- 1) Novel NOV1a (SEQ ID NO:2)
- 2) Novel NOV1b (SEQ ID NO:4)
- 3) Novel NOV1c (SEQ ID NO:6)
- 4) Novel NOV1d (SEQ ID NO:8)
- 4) $\frac{gi|13569928|ref|NP-112217.1|}{gi|13569928|NP 112217}$ a disintegrin-like and metalloprotease with thrombospondin type 1 motif, 12 [Homo sapiens] (SEQ ID NO:30)
- 5) $\frac{gi|11493589|gb|AAC35563.1|AF163762.1}{gi|11493589}$ $\frac{AAG35563}{AAG35563}$ (AF163762) zinc metalloendopeptidase [Homo sapiens] (SEQ ID NO:31)
- 6) gi|10645199|ref|NP_055087.1| gi|10645199| NP_055087 a disintegrin and
 metalloprotease with thrombospondin motifs-7 preproprotein; a disintegrin-like and
 metalloprotease (reprolysin type) with thrombospondin type 1 motif, 7 [Homo sapiens]
 (SEQ ID NO:32)
- 7) gi|15309931|ref|XP 054419.1| gi|15309931| XP 054419 a disintegrin-like and metalloprotease domain with thrombospondin type I repeats 10 [Homo sapiens] (SEQ ID NO:33)
- 8) gi|7656869|ref|NP 055088.1| gi|7656869| NP 055088 a disintegrin and metalloprotease with thrombospondin motifs-6 preproprotein; a disintegrin-like and metalloprotease (reprolysin type) with thrombospondin type 1 motif, 6 [Homo sapiens] (SEQ ID NO:34)

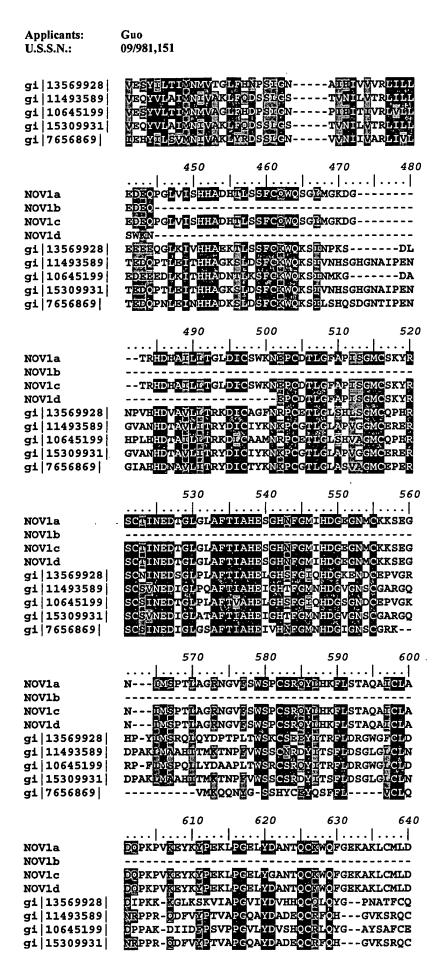
Applicants: U.S.S.N.:

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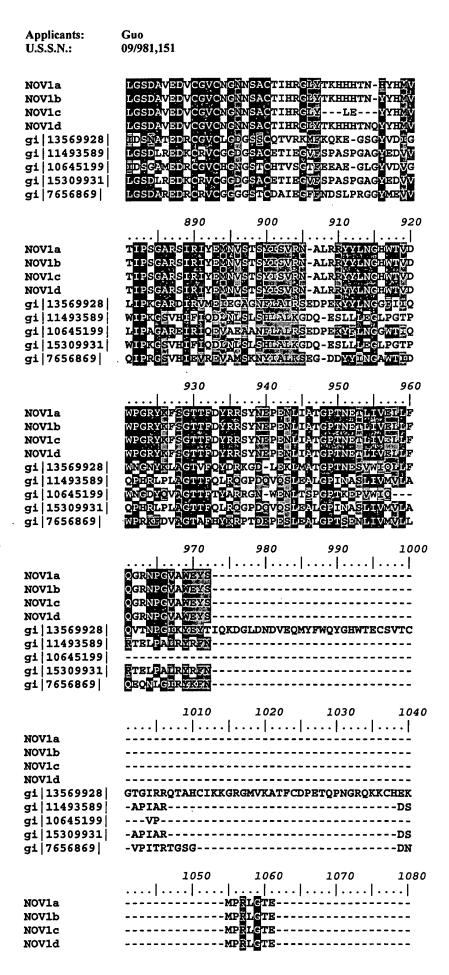
| | 10 20 30 40 |
|-----------------------------|---|
| | |
| NOV1a | MKPRARGWRGLAALWMLLAQVAEQVSPGRSHQRGNRGSGQ |
| NOV1b | MKPRARGWRGLAALWMLLAQVAEQVSPGRSHQRGNRGSGQ |
| NOV1c | |
| NOV1d | MKPRARGWRGLAALWMLLAQVAEQVSPGRSHQRGNRGSGQ |
| gi 13569928 | |
| gi 11493589 | |
| gi 10645199 | |
| gi 15309931 | |
| gi 7656869 | |
| | |
| | 50 60 70 80 |
| | |
| NOV1a | LEASPPRLLSRGPRRLTAMSPLFSAGTCVRHGTRSGSAWE |
| NOV1b | LEASPPRLLSRGPRRLTAMSPLFSAGTCVRHGTRSGSAWE |
| NOV1c | GSPSV |
| NOV1d | LEASPPRLLSRGPRRLTAMSPLFSAGTCVRHGTRSGSAWE |
| gi 13569928 | MPCAQRSWLANLSVVAQLLNFGALCYGRQPQPGPVRF |
| gi 11493589 | |
| gi 10645199 | MPGGPSPRSPAPLLRPLLLLLCALAPG-APGPAPGRA |
| gi 15309931 | |
| gi 7656869 | MEILWKTLTWILSLIMASSEFHSDHRL |
| | |
| | 90 100 110 120 |
| NOV1a | PERPASSITGAAGLDGKGRDMDEAGNHRSOOTNTGTENO |
| NOV1a NOV1b | PERPASSIRGAAGLDGKGRDMDEAGNHRSQQINIGIENQ PERPASSSTRGAAGLDGKGRDMDEAGNHRSQQINIGIENQ |
| NOV16 | PRPPP |
| NOV1d | PERPASSSTRGAAGLDGKGRDMDEAGNHRSOOTNTGTENO |
| gi 13569928 | PDRROEHFIKGL |
| gi 11493589 | RSQDEFLSSL |
| gi 10645199 | TRCR |
| gi 15309931 | |
| gi 7656869 | SYSSOEEFLTYL |
| 3-1.000001 | |
| | 130 140 150 160 |
| | |
| NOV1a | TLHVLT-QYDIVSAYEVDHRCDYVSHE MHHQR-RRRAVA |
| NOV1b | TLHVLTREYDEVSAYEVDHRGDXVSHE MHHOR-RRAVA |
| NOV1c | DLVSAYEVDHRGDYVSEEFIMHHOR-RRRAVA |
| NOV1d | TLHVLT-QYDIVSAYEVDHRGDYVSHEEMHHQR-RRRAVA |
| gi 13569928 | PEYHÜVGPVRVDASGHERESYGEHYPITSSRRKRD |
| gi 11493589 | es <u>yet</u> afptr <mark>vd</mark> hngal <u>ka</u> fspppp-rro <u>rr</u> gtg |
| gi 10645199 | AALDEVHPVRVDAGGSEESYEEWPR-ALRKRD |
| g1 15309931 | |
| gi 7656869 | ehyogtipirydongasesftyknd-khsrrrrs |
| | |
| | 170 180 190 200 |
| | |
| NOV1a | VSEVES-THLRTKGPRHDFHMDTRTSSSLVAPGT |
| NOV1b | VSBVES-CHLRUKGPRHDFHMDLRTSSSLVAPGFI |
| NOV1c | VSevep-aflöycrarelrlcveafpiansopggi VSeves-ühlrikgprhofhmodrtssslyapgei |
| NOV1d | |
| gi 13569928 | edgsedw-wyyrisheerdlffutviogfesns <u>yi</u> aiaesrufykvaspsthfllautresrliaghvs |
| gi 11493589 | VSVRRDAPAFYELQYRGRELRFALTANQHLLAPGEV |
| gi 10645199 gi 15309931 | APAVENTENT TENT TOUR TOUR THE |
| gi 15309931 gi 7656869 | Mdpidpqqavskiffkisaygkhfhlnitlntdfyskhft |
| 97 1020003 | Spr 121 XXV - Number : William Carte market 1 12 12 1 12 12 12 12 12 12 12 12 12 12 |
| | 210 220 230 240 |
| | |

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|--|--|
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | VOTLGETETKSVQTLPP-EDFCFYQGSLESHRNSPSHG VOTLGETETKSVQTLPP-EDFCFYQGSLESHRNSPSHG NLSNVRSHWREQHASKRII-TNAMLGESALASTRKSN VOTLGETETKSVQTLPP-EDFCFYQGSLESHRNSPSHG MEKR-YGNLSHVKMMASSAPLCHLSGTVLQQGTRVGTA VEYWTEGLAWQRAARPHCLYAGHIGGQASSS-HV SETRREGGLGRAHIRAHTPACHLLGEVQDPELEGGLA VEYWGEDGPQWKHDFLDNCHYTGYLQDQRSET-KV |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 250 260 270 280 GKFCEGSTRTLKLCNSQKCPRDSVDFRAAQCAEHNSRRFR GKFCEGSTRTLKLCNSQKCPRDSVDFRAAQCAEHNSRRFR CVFFL-SFYFFQSGMIRTEEADYFLRPLPSHLSWKLG GKFCEGSTRTLKLCNSQKCPRDSVDFRAAQCAEHNSRRFR ALSACHGLTGFFQLPHGDFFIEPVKKHPLVEG- AISTCGGLHGLIVADEESYLIEPLHGGPKGS AISACDGLKGVFQLSNEDYFIEPLDSAPARPG- ALSNCVGLHGVIATEDEEXFIEPLKNTTEDSKH |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | GRHYKWKPTTQVEXDLCKLYCHAEGFDFFFSLENKTRDGT GRHYKWKPTTQVEQDLCKLYCHAEGFDFFFSLENKTRDGT RAAQGSSESHVLYKREVLVTSRTWELAHQPLHESDLRLGL GRHYKWKPTTQVEADLCKLYCHAEGFDFFFSLENKTRDGTGYHPHIVYRRQKVPETKEP-ECGENDSV RSPEESGPTVVYRRSSLRHPHIDTACGV-RDEKFWKGRPWHAQPHVVYRRQAPER-LAQRGDSSAPSECGYQVYP FSYENGHPHVIYRKSALQQRHEYDHSHCGVSDFTRSGKPW |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 330 340 350 360 PCSEDSRNVCIDGICELSVVSTSAHMPQPPKEDLFILPDE PCSEDSRNVCIDGICE |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 370 380 390 400 YESCLERKESILESHENE LNVETLVVVDKKMÖNHEHEN YESCLERKESILESHENE LNVETLVVVDKKMÖNHEHEN YESCLERKESILESHENE LNVETLVVVDKKMÖNHEHEN YESCLERKESILESHENE LNVETLVVVDKKMÖNHEHEN YESCLERKESILESHENE LNVETLVVDKKMÖNHEHEN WERHNLPSESISESERSISE RWVETLVVADTKMETHESEN LGNETERG PGLERSVSE RYVETLVVADKMMATHERED QÖWRRPELER LHORSVSE KWVETLVVADAKMVETHE QPQ MMVATHERED LPINNTHIHHROKESVSI BEFVETLVVADKMWOGYHERED |
| NOV1a NOV1b NOV1c NOV1d | 410 420 430 440 |

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|--|--|
| gi 7656869 | skxнно |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | FKKDICKALWCHRIGRKCETKFMPAAEGTICGHDMWDICKALWCHRIGRKCETKFMPAAEGTICGHDMW FKKDICKALWCHRIGRKCETKFMPAAEGTICGHDMW FKKDICKALWCHRIGRKCETKFMPAAEGTICGHDMW FKKDICKALWCHRIGRKCETKFMPAAEGTICGHDMW EVENVCOTLWCSVKG-FCRSKLDAAADGTOCGEKKW KYGEVCSELWCLSKSNECITNSEPAAEGTICOTHTIDKGW DMDNVCHTLWCSVGT-TCHSKLDAAVDGTREGENKW KYGEVCSELWCLSKSNECITNSEPAAEGTICOTHTIDKGW LFREVCRELWCLSKSNECVTNSEPAAEGTICOTHTIDKGW |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 690 700 710 720 |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 730 740 750 760 RSR-LCTNPNPSHGGKECEGSTRTLKLCNSOKCPRDSVDF RSR-LCTNPRPSHGGKECEGSTRTLKLCNSOKCPRDSVDF RSRSONTHSRPSHGGKECEGSTRTLKLCNSOKCPRDSVDF RSR-LCTNPRPSHGGKECEGSTRTLKLCNSOKCPRDSVDF RSR-LCTNPRPSHGGKECEGSTRTLKLCNSOKCPRDSVDF SSR-LCNNPEPKFGGKYCTGERKRYRLCNVHPCRSEPTF SSR-HCDSPRPHIGGKYCTGERKRYRLCNVHPCRSEPTF SSR-HCDSPRPHIGGKYCTGERKRYRLCNLOACPAGRPSF SSR-HCDSPRPHIGGKYCTGERKRHRSCNTDDCPPGSQDF SIR-HCDSPRPHIGGKYCTGERKRYRSCNTDDCPPGSQDF SIR-HCDSPRPSGGGKYCTGERKRYRSCNTDPCPLGSRDF |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 770 780 790 800 RAAO CAEHNSRRFRGREYKWKP |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 810 820 830 840 FDFFFSSNKVKDGTPCSEDSRNVCIDGICEX-GCDNV FDFFFSSNKVKDGTPCSEDSRNVCIDGICEGCDNV FDFFFSSNKVKDGTPCSEDSRNVCIDGICERVGCDNV FDFFFSSNKVKDGTPCSEDSRNVCIDGICERVGCDNV GOFSEKMLDAVIDGTPCFEGGNSRNVCINGICKMVGCDYE FNFYTERAAAVVDGTPCRPDEVDICVSGECKHVGCDRV EYFAKKTRDAVVDGTPCVQVRASROTCINGICKNVGCDFE FNFYTERAAAVVDGTPCRPDEVDICVSGECKHVGCDRV WNFYTERAPAVIDGTQCNADSLETCINGECKHVGCDRV |
| | |



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|--|---|
| gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | ACPPRWWAGEWEACSATC PHGEKKRTVLCIQTMVSDEQA LPPYSWHYAPWTKCSA CAGGSQV-QAVECRNQLDSSAASR PGGGS-RGG LPPYSWHYAPWTKCSA CAGGSQV-QAVECRNQLDSSA EVGFTWNHQPWSECSATCAGG |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1090 1100 1110 1120 . |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1130 1140 1150 1160 |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1170 1180 1190 1200 |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1210 1220 1230 1240 |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 | 1250 1260 1270 1280 |

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|--|---|
| gi 7656869 | KET <mark>LL</mark> |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1290 1300 1310 1320 |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1330 1340 1350 1360 |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1370 1380 1390 1400 PPLTPDLSRESWWPPFSTVMEGLLPSQRPTTSETGTPRVE |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1410 1420 1430 1440 GMVTEKPANTLLPLGGDHQPEPSGKTANRNHLKLPNNMNQ |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1450 1460 1470 1480 TKSSEPVLTEEDATSLITEGFLLNASNYKQLTNGHGSAHWPVFS |

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| | · |
| NOV1a | |
| NOV1b | |
| NOV1c | |
| NOV1d | TYPOTH COMMON OF THE PROPERTY |
| gi 13569928 gi 11493589 | IVGNWSECSTTCGLGAYWKRVECTTQMDSDCAAIQRPDPA VAGEWGECSAQCGVGQRQRSVRCTS |
| gi 10645199 | HYGPWTKCTVTCGRGE |
| gi 15309931 | VAGEWGECSAQCGVGQRQRSVRCTS |
| gi 7656869 | |
| | 1530 1540 1550 1560 |
| | |
| NOV1a | |
| NOV1b | |
| NOV1c NOV1d | |
| gi 13569928 | KRCHLRPCAGWKVGNWSKCSRNCSGGFKIREIQCVDSRDH |
| gi 11493589 | HTGQASHECTEALR |
| gi 10645199 | LVSGQGH |
| gi 15309931 gi 7656869 | HTGQASHECTEALR |
| 91,7030007 | |
| | 1570 1580 1590 1600 |
| | |
| NOV1a NOV1b | |
| NOV1c | |
| NOV1d | |
| gi 13569928 | RNLRPFHCQFLAGIPPPLSMSCNPEPCEAWQVEPWSQCSR |
| gi 11493589 gi 10645199 | |
| gi 15309931 | PPTTQQCEA |
| gi 7656869 | |
| | 1610 1620 1630 1640 |
| | |
| NOV1a | |
| NOV1b | |
| NOV1c NOV1d | |
| gi 13569928 | SCGGGVQERGVFCPGGLCDWTKRPTSTMSCNEHLCCHWAT |
| gi 11493589 | KCDSPTPG |
| gi 10645199 gi 15309931 | HCWAT |
| gi 7656869 | |
| - ' ' | |
| | 1650 1660 1670 1680 |
| NOV1a | |
| NOV1b | |
| NOV1c | |
| NOV1d | |
| gi 13569928 gi 11493589 | GNWDLCSTSCGGGFQKRIVQCVPSEGNKTEDQDQCLCDHK DGPEECKDVN |
| gi 10645199 | TGLEVCFSEE |
| gi 15309931 | DGPEECKDVN |
| gi 7656869 | |
| | 1690 1700 1710 1720 |
| | |
| NOVla | |
| NOV1b | |
| NOV1c NOV1d | |
| | • |

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|--|---|
| gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | PRPPEFKKCNQQACKKSADLLCTKDKLSASFCQTLKAMKKKVAYCPLVLKFQFPQFS-ICEMRLAIALKVAYCPLVLKFQF |
| NOV1a NOV1b NOV1c NOV1d gi 13569928 gi 11493589 gi 10645199 gi 15309931 gi 7656869 | 1730 1740 1750 |

At page 32, please replace paragraph at lines 31-42 with the following new paragraph:

The presence of identifiable domains in NOV1, as well as all other NOVX proteins, was determined by searches using software algorithms such as PROSITE, DOMAIN, Blocks, Pfam, ProDomain, and Prints, and then determining the Interpro number by crossing the domain match (or numbers) using the European Bioinformatics Institute Interpro. website (http://www.ebi.ac.uk/ interpro). DOMAIN results for NOV1 as disclosed in Tables 1L-IO, were collected from the Conserved Domain Database (CDD) with Reverse Position Specific BLAST analyses. This BLAST analysis software samples domains found in the Smart and Pfam collections. For Table 1E and all successive DOMAIN sequence alignments, fully conserved single residues are indicated by black shading or by the sign (|) and "strong" semi-conserved residues are indicated by grey shading or by the sign (+). The "strong" group of conserved amino acid residues may be any one of the following groups of amino acids: STA, NEQK (SEQ ID NO: 161), NHQK (SEQ ID NO: 162), NDEQ (SEQ ID NO: 163), QHRK (SEQ ID NO: 164), MILV (SEQ ID NO: 165), MILF (SEQ ID NO: 166), HY, FYW.

At page 33, please replace Table 1L with the following:

Table 1L. Domain Analysis of NOV1

gnl|Pfam|pfam01421, Reprolysin, Reprolysin (M12B) family zinc metalloprotease. The members of this family are enzymes that cleave peptides. These proteases require zinc for catalysis. Members of this family are also known as adamalysins. Most members of this family are snake venom endopeptidases, but there are also some mammalian proteins, and fertilin. Fertilin and closely related proteins appear to not have some active site residues and may not be active enzymes. (SEQ ID NO:76)

CD-Length = 199 residues, 95.0% aligned Score = 122 bits (307), Expect = 7e-29

```
Query:
       378
            DKKMMONHG--HENITTYVLTILNMVSALFKDGTIGGNINIAIVGLILLEDEOPGLVISH
                             | |+|+|+ +++
                                                 || + +||| +
                   +
                           l
            DHGMFTKYGSDLNKIRQRVHQIVNLVNEIYRPL----NIRVVLVGLEIWSDGDK-ITVQG
Sbjct:
       10
Query:
       436
            HADHTLSSFCQWQSGLMGKDGTRHDHAILLTGLDICSWKNEPCDTLGFAPISGMCSKYRS
                                   ||+| ||||+|
             |+ || |+|+
                            +
                                                       +|+| | + ||||
            DANDTLHRFLEWRETDLLKR-KSHDNAQLLTGIDF-----DGNTIGAAYVGGMCSPKRS
Sbjct:
       65
                                                                          117
            CTINED -- TGLGLAFTIAHESGHNFGMIHDGEGN-MCKKSEGNIMSPTLAGRNGVFSWS
Query:
       496
                                                                          551
                       | +| |+||| ||| |||
              + +|
                                                 +
                                                      | ||+| +
            VGVVQDHSPIVLLVAVTMAHELGHNLGMTHDDINKCTCEGGGGCIMNPVASSSPGKK-FS
Sbjct:
       118
            PCSRQYLHKFLSTAQAICLADQ 573
Query:
                                         (aa 378-573 of SEQ ID NO:2)
                    +||+ + || ++
             11
       177 NCSMDDYQQFLTKGKPQCLLNK 198
                                         (SEQ ID NO:76)
Sbjct:
```

At page 33, please replace Table 1M with the following:

Table 1M. Domain Analysis of NOV1 gnl|Smart|smart00209, TSP1, Thrombospondin type 1 repeats; Type 1 repeats in thrombospondin-1 bind and activate TGF-beta. (SEQ ID NO:77) CD-Length = 51 residues, 100.0% aligned Score = 63.5 bits (153), Expect = 5e-11

```
        Query:
        668
        WSDWSSWSPCSRTCGGGVSHRSRLCTNPNPSHGGKFCEGSTRTLKLCNSQKCP
        720 (of SEQ ID

        NO:2)
        Sbjct:
        1
        WGEWSEWSPCSVTCGGGVQTRTRCCNPPPN--GGGPCTGPDTETRACNEQPCP
        51 (SEQ ID NO:77)
```

At page 33, please replace Table 1N with the following:

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Guo 09/981,151

Table 1N. Domain Analysis of NOV1

gnl|Pfam|pfam00090, tsp_1, Thrombospondin type 1 domain. (SEQ ID
NO:78)
CD-Length = 48 residues, 100.0% aligned
Score = 54.7 bits (130), Expect = 2e-08

Query: 669 SDWSSWSPCSRTCGGGVSHRSRLCTNPNPSHGGKFCEGSTRTLKLCNSQKC 719 (of SEQ ID NO:2)

| | | | | | | | | + | | | + + | |

Sbjct: 1 SPWSEWSPCSVTCGKGIRTRQRTCNSP---AGGKPCTGDAQETEACMMDPC 48 (SEQ ID NO:78)

At page 34, please replace Table 10 with the following:

Table 10. Domain Analysis of NOV1

gnl|Pfam|pfam01562, Pep_M12B_propep, Reprolysin family propeptide.
This region is the propeptide for members of peptidase family M12B.
The propeptide contains a sequence motif similar to the "cysteine switch" of the matrixins. This motif is found at the C terminus of the alignment but is not well aligned. repeats. (SEQ ID NO:79)
CD-Length = 117 residues, only 39.3% aligned
Score = 38.5 bits (88), Expect = 0.002

Query: 180 LRTSSSLVAPGFIVQTLGKTGTKSVQTLPPEDFCFYQGSLRSHRNS 225 (of SEQ ID NO:2)

| + ||+|| | | | + + ||+||| + + ||
Sbjct: 2 LEKNRSLLAPDFTVTTYDDDGTLVTEHPLIQDHCYYQGYVEGYPNS 47 (SEQ ID NO:79)

At page 41 please replace Table 2C with the following:

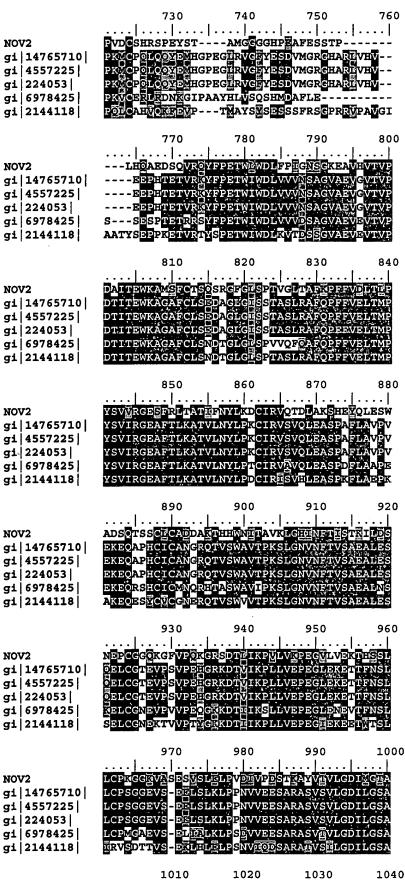
| Table 2C. BLAST results for NOV2 | | | | | |
|---|--|--------|-------------------|-------------------|--------|
| Gene Index/ | Protein/ | Length | Identity | Positives | Expect |
| Identifier | Organism | (aa) | (%) | (%) | |
| gi 14765710 ref XP 006925.4 XP 006925 | alpha 2 macroglobulin precursor [Homo sapiens] | 1474 | 593/1486 (39%) | 870/1486 (57%) | 0.0 |
| <u>gi 4557225 ref NP_0</u> <u>00005.1 </u> NP_000005 | alpha 2 macroglobulin precursor [Homo sapiens] | 1474 | 591/1486 (39%) | 869/1486 (57%) | 0.0 |
| gi 224053 prf 1009 174A 1009174A | macroglobulin alpha2 [Homo sapiens] | 1450 | 585/1471 (39%) | 861/1471 (57%) | 0.0 |

| gi 6978425 ref NP 0 36620.1 NP 036620 | alpha-2- macroglobulin [Rattus norvegicus] | 1472 | 578/1483 (38%) | 867/1483 (57%) | 0.0 |
|---|--|------|-------------------|-------------------|-----|
| gi 2144118 pir JC5 143 JC5143 | alpha- macroglobulin precursor - guinea pig | 1476 | 570/1495 (38%) | 858/1495 (57%) | 0.0 |

At page 41 please replace Table 2D with the following:

Table 2D. ClustalW Analysis of NOV2

- NOV2 (SEQ ID NO:10) gi | 14765710 | ref | XP 006925.4 | gi | 14765710 | XP 006925 alpha 2 macroglobulin precursor [Homo sapiens] (SEQ ID NO:35) gi|4557225|ref|NP 000005.1| gi|4557225| NP 000005 alpha 2 macroglobulin precursor [Homo sapiens] (SEQ ID NO:36) gi|224053|prf||1009174A gi|224053| 1009174A macroglobulin alpha2 [Homo sapiens] 3) (SEQ ID NO:37) gi|6978425|ref|NP 036620.1| gi|6978425| NP 036620 alpha-2-macroglobulin [Rattus 4) norvegicus] (SEQ ID NO:38) gi|2144118|pir||JC5143 gi|2144118| JC5143 alpha-macroglobulin precursor - guinea pig (SEQ ID NO:39)
- 590 600 570 580 NOV2 gi | 14765710 | RLLIYAVLENGOVIGDSAKYOVENGLANKVOLSESE RLLIYAVLENGOVIGDSAKYOVENGLANKVOLSESE RLVZYATLENGOVIJGOBAKYOVENCLONKVOLVERE RLVZYATLENGOVIJGOBAKYOVENCLONKVOLSESE VIGDSAKY DVENCLANKYDLSFSP gi | 4557225 | gi | 224053 | gi | 6978425 | R<mark>uliya lpsgria</mark>dsaky<u>n</u>vencl<mark>d</mark>nkviilsfs<mark>e</mark> gi | 2144118 | NOV2 **CALRAVDISSVLLIPRIPDISE** PASHAHLRVTÄAPQS PASHAHLRVTÄAPQS CALRAVDOSVLLMKPDAELSAS gi | 14765710 | CALRAVDOSVLLMKPDAELSASSV gi | 4557225 | CALRAVDOSVĽLMKPDAEĽSA Pas<mark>hahlrvtaaposy</mark> Pa<mark>irailsvm</mark>asposi gi | 224053 | CCLRAVDOSVLLMKP gi | 6978425 | THLRVTASPOST CALRAVDOSVLLRKPE gi | 2144118 | 650 660 670 NOV2 gi | 14765710 | gi | 4557225 | gi | 224053 | gi | 6978425 | vkoltgfpogadoreedtng--YALLPVKDLTGFPGLLGOOBENDGE----gi | 2144118 | 690 700 710 720 GESSORSIINRPSFSEG-TOMESFFRDVGLKII NOV2 YINGI<mark>TYEPVS</mark>STNEKDMYSFLEDMGLK gi | 14765710 | gi | 4557225 | ATINGI<mark>T</mark>ASABEBÄ<mark>I</mark>NEKDWASŁIKOWGTKAŁINSKISK ATINGITARAN ZINEKDWASŁIKOWGTKAŁINSKISK ATINGITARAN ZINEKOWASŁIKOWGTKAŁINSKISK ATINGITARAN ZINEKOWASŁIKOWGTKAŁINSKISK gi | 224053 | gi | 6978425 | gi | 2144118 |



| U.S.S.N.: | 09/981,151 |
|--|--|
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | TONLOGLEOMPSGCGEONMVLFAPITYVLOVLETAGLLTE MONTONLLOMPYGCGEONMVLFAPNIYVLDYLNETQOLTP MONTONLLOMPYGCGEONMVLFAPNIYVLDYLNETQOLTP MONTONLLOMPYGCGEONMVLFAPNIYVLDYLNETQOLTP MONTONLLOMPYGCGEONMVLFAPNIYVLDYLNETQOLTO MONTONLLOMPYGCGEONMVLFAPNIYVLDYLNETQOLTO |
| | 1050 1060 1070 1080 |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | EIRSRAYGELETGYQEELMYKHSYGSYSAFGERDG NGN EIKSKAIGYLNTGYQRQLNYKHYDGSYSTFGERYGRNQGN EIKSKAIGYLNTGYQRQLNYKHYDGSYSTFGERYGRNQGN EIKSKAIGYLNTGYQRQLNYKHYDGSYSTFGERYGRNQGN EIKEKAIAYLNTGYQRQLNYKHRDGSYSTFGENYRGGQGN EIKSKAIAYLNTGYQRQLNYKHRDGSYSTFGENYRGGQGN |
| | 1090 1100 1110 1120 |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | TWLTAFVIKCFGQAOKGIFIDPKNIODALWWW.GNOLPSG TWLTAFVLKTFAOARAYIFIDEAHITOALIWESOROKDNG TWLTAFVLKTFAOARAYIFIDEAHITOALIWESOROKDNG TWLTAFVLKTFAOARAYIFIDEAHITOALIWESOROKDNG TWLTAFVLKSFAOARAYIFIDEAHITOALIWESOOKDNG TWLTAFVLKSFAOARAYIFIDEAHITOALIWESOOKDNG TWLTAFVLKTFSOARAYIFIDEAHITOALIWESOSOKDNG |
| | 1130 1140 1150 1160 |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | CVANVGNLLETANKGGVDDEVSLEAVUTAALLENGKDVDD CFRSSGSLLNNAIKGGVEDEVTLSAYITIALLENGKDVDD CFRSSGSLLNNAIKGGVEDEVTLSAYITIALLENDLTVTH CFRSSGSLLNNAIKGGVEDEVTLSAYIKIALLENDLTVTH CFRSSGSLLNNAIKGGVEDEVTLSAYITIALLENSLPVTH CFWSSGSLLNNAIKGGVEDE |
| | 1170 1180 1190 1200 |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | PZVSQGLRCLENSATET |
| | 1210 1220 1230 1240 |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | EMDIRNIILKOIDOOAIISGESTYMSÖKPTPSINASPISE NODERKEULKSINEEAVKEDNSVHWERPOKPKAPVGHEYE NODERKEULKSINEEAVKEDNSVHWERPOKPKAPVGHEYE NODERKEULKSINEEAVKEDNSVHWERPOKPKAPVGHEYE NODERKEILKSINEEAVKEISVHWTRPOKPSVSVGLWYO NOERKEILKSIEDEGVKEDNSIHWARPOKPKVSEGPLEK |
| | 1250 1260 1270 1280 |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | PAAVOVEETAYALLAOLT- PELTO GETAKATSIVAWI POAPSAEVEMTSYVLLAYLTA PAPTSEDLTSATAIVKWE POAPSAEVEMTSYVLLAYLTA PAPTSEDLTSATAIVKWE POAPSAEVEMTSYVLLAYLTA PAPTSEDLTSATAIVKWE POATSAEVEMTÄYVLLAYLTA PAPTOEDLTAMLIYKWE SOAPSAEVEMTÄYVLLAYLTA PAPTOEDLTAMLIYKWE |
| | 1290 1300 1310 1320 |

Applicants:







| Applicants: U.S.S.N.: | Guo 09/981,151 |
|--|--|
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | AKOHNAYGGFSSTQDTVVALQALEKYATTAYMPS EBINE TKQQNAQGGFSSTQDTVVALHALSKYGAATFTRT KAAQV TKQQNAQGGFSSTQDTVVALHALSKYGAATFTRT KAAQV TKQQNAQGGFSSTQDTVVALHALSKYGAATFTRT KAAQV TKQQNEHGGFSSTQDTVVALHALSKYGEATFTRT KAAQV TKQQNEHGGESSTQDTVVALHALSKYAAATFTRT KAAQV |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | 1330 1340 1350 1360 |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | 1370 1380 1390 1400 |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | 1410 1420 1430 1440 RSLTEGERTSYVGSRSSSNMAIVEVKMESGFSPEGTNOT TSFQISLSVSYTGSRSSSNMAIVDVKMVSGFIPLKPTVKM TSFQISLSVSYTGSRSSSNMAIVDVKMVSGFIPLKPTVKM TSFQISLSVSYTGSRSSSNMAIVDVKMVSGFIPLKPTVKM TSFQISLSVSYTGSRSSSNMAIVDVKMVSGFIPLKPTVKM TSFQISLSVSYTGSRSSSNMAIADVKMVSGFIPLKPTVKM TSFQISLSVSYTGSRSSSNMAIADVKMVSGFIPLKPTVKM |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | 1450 1460 1470 1480 LECOPLYKKVEFGTTTNIYLDETIKNTOTYTTTESOSYL LERSNHYSKTEVSSNHYLIYLDKYSNOTUSIFFTYPODYP LERSNHYSKTEVSSNHYLIYLDKYSNOTUSIFFTYPODYP LERSNHYSKTEVSSNHYLIYLDKYSNOTUSIFFTYPODYP LERSVHYSKTEVSSNHYLIYLDKYSNOTUSIFFTYPODYP LERSVHYSKTEVSNNHYLIYLDKYSNOTUSIFFTYPODTE LEGSEHTSKTEVSNNHYLIYLDKYSNOTUSISFTYOODTE |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | 1490 1500 1510 1520 VINLKPATIKVYDYYLPGSFKIRSOYTIVWSMINDSIVDSV VRDLKPAIVKVYDYYET GEFAIAEY APCSKOLIGNA VRDLKPAIVKVYDYYET DEFAIAEY APCSKOLIGNA VRDLKPAIVKVYDYYET DEFAIAEY APCSKOLIGNA GROLKPAIVKVYDYYEKDEFAIAEY APCSKOLIGNA VRDLKPAIKVYDYYEKDEFAIAEY APCSKOLIGNA VRDLKPAIKVYDYYEKDEFAIAEY APCSKOLIGNA |
| NOV2 gi 14765710 gi 4557225 gi 224053 gi 6978425 gi 2144118 | 1530 1540 ARHPEPPPFKTEAFIPSLPGSVNN (SEQ ID NO:10) (SEQ ID NO:35) (SEQ ID NO:36) (SEQ ID NO:37) (SEQ ID NO:38) (SEQ ID NO:39) |



Pages 45, please replace Table 2E with the following

Table 2E Domain Analysis of NOV2

gnl|Pfam|pfam00207, A2M, Alpha-2-macroglobulin family. This family
includes the C-terminal region of the alpha-2-macroglobulin family.
(SEQ ID NO:80)
CD-Length = 751 residues 98.5% aligned

CD-Length = 751 residues, 98.5% aligned Score = 563 bits (1451), Expect = 2e-161

| Query: | 728 | EDSQVRQYFPETWLWDLFPIGNSGKEAVHVTVPDAITEWKAMSFCTSQSRGFGLSPTVGL | 787 |
|--------|------|--|------|
| Sbjct: | 4 | + + + ++ + ++ + + + ++ ++ ++ DDITIRSYFPESWLWEVEEVDRSPVLTVNITLPDSITTWEILAVSLSNTKGLCVADPVEL | 63 |
| Query: | 788 | TAFKPFFVDLTLPYSVVRGESFRLTATIFNYL-KDCIRVQTDLAKSHEYQLESWADSQTS | 846 |
| Sbjct: | 64 | + ++ ++ | 115 |
| Query: | 847 | SCLCADDAKTHHWNITAVKLGHINFTISTKILDSNEPCGGQKGFVPQKGRSDTLIKPVLV | 906 |
| Sbjct: | 116 | FCSLATORTRSSQSVRPKSLSSVSFPVVVVPLASGLSLVEVVASVPEFFVKDAVVKTLKV | 175 |
| Query: | 907 | KPEGVLVEKTHSSLLCPKGGKVASESVSLELPVDIVPD-STKAYVTVLGDIMGTALQ + | 962 |
| Sbjct: | 176 | EPEGARKEETVSSLLLPPEHLGGGLEVSEVPALKLPDDVPDTEAEAVISVQGDPVAQAIQ | 235 |
| Query: | 963 | NEIRSRAVGFLEIG | 1013 |
| Sbjct: | 236 | + +++ + + ++ + + + + | 295 |
| Query: | 1014 | YQKELMYKHSNGSYSAFGERDGNGNTWLTAFVTKCFGQAQKFIFIDPKNIQDALKW-MAG | 1072 |
| Sbjct: | 296 | ++ + ++ + + + ++ ++ + + + + + + + + + | 353 |
| Query: | 1073 | NQLPSGCYANVGNLLHTAMKGGVDDEVSLTAYVTAALLEMGKDVDDPMVSQGLRCL | 1128 |
| Sbjct: | 354 | QQKDDGVFRESGPVIHNEMKGGVGDDAEVEVTLTAFITIALLEAKLVCISPVVANALSIL | 413 |
| Query: | 1129 | KNSATSTTNLYTQALLAYIFSLAGEMDIRNILLKQLDQQAIISGESIYWSQK | 1180 |
| Sbjct: | 414 | KASDYLLENYANGQRVYTLALTAYALALAGVLHKLKEILKSLKEELYKALVKGHWERPQK | 473 |
| Query: | 1181 | PTPSSNASPWSEPAAVDVELTAYALLAQLTKPSLTQKEIAKATSIVAWLAKQHNAYGGFS | 1240 |
| Sbjct: | 474 | + + + + + + + | 531 |
| Query: | 1241 | STQDTVVALQALAKYATTAYMPSE-EINLVVKSTEN-FQRTFNIQSVNRLVFQQDTLP-N | 1297 |
| Sbjct: | 532 | + + +++ ++ + + + + STQDTVMALQALSKYGIATPTHKEKNLSVTIQSPSGSFKSHFQILNNNAFLLRPVELPLN | 591 |
| Query: | 1298 | VPGMYTLEASGQGCVYVQTVLRYNILPPTNMKTFSLSVEIGKARCEQPTSPR-SLTLTIH | 1356 |
| Sbjct: | 592 | EGFTVTAKVTGQGTLTLVTTYRYKVLDKKNTFCFDLKIETVPDTCVEPKGAKNSDYLSIC | 651 |
| Query: | 1357 | TSYVGSRSSSNMAIVEVKMLSGFSPMEGTNQLLLQQPLVKKVEFGTDTLNIYLDELIK | 1414 |
| Sbjct: | 652 | TRYAGSRSDSGMAIADISMLTGFIPLKPDLKKLENGVDRYVSKYEIDGNHVLLYLDKVSH | 711 |
| Query: | 1415 | -NTQTYTFTISQSVLVTNLKPATIKVYDYYLP 1445 (of SEQ ID NO:10) | |

Sbjct: 712 SETECVGFKIHQDFEVGLLQPASVKVYDYYEP 743 (SEQ ID NO:80)

Pages 46, please replace Table 2F with the following:

Table 2F Domain Analysis of NOV2

gnl|Pfam|pfam01835, A2M_N, Alpha-2-macroglobulin family N-terminal
region. This family includes the N-terminal region of the alpha-2macroglobulin family. (SEQ ID NO:81)
CD-Length = 620 residues, 98.4% aligned
Score = 236 bits (603), Expect = 5e-63

| Query: | 5 | LLLGMLALSPAIAEELPNYLVTLPARLNFPSVQKVCLDLSPGYSDVKFTVTLETKDKT | 62 |
|--------|-----|---|----------|
| Sbjct: | 2 | + | 61 |
| Query: | 63 | QKLLEYSGLKKRHLHCISFLVPPPAGGTEEVATIRVSGVGNNISFEEKKKVLIQ + | 116 |
| Sbjct: | 62 | + + + + + + + + RNLSSLFTVLLSSKDLFHCVSFTVPQPGLFKSSKGEESFVVVQVKGPTHTFKEKVTVLVS | 121 |
| Query: | 117 | RQGNGTFVQTDKPLYTPGQQVYFRIVTMDSNFVPVNDKYSMVELQDPNSNRIAQWLEVVP + + + + ++ ++ +++ ++ + ++ + | 176 |
| Sbjct: | 122 | SRRGLVFIQTDKPIYTPGQTVRYRVFSVDENLRPLNELI-LVYIEDPEGNRVDQWEVNKL | 180 |
| Query: | 177 | EQGIVDLSFQLAPEAMLGTYTVAVAEGKTFGTFSVEEYVLSPFLLLLSSVLPKFK | 231 |
| Sbjct: | 181 | EGGIFQLSFPIPSEPIQGTWKIVARYESGPESNYTHYFEVKEYVLPSFEVS | 231 |
| Query: | 232 | VEVVEPKELSTVQESFLVKICCRYTYGKPMLGAVQVSVCQKANTYWYREVEREQLPDKCR + + + +++ | 291 |
| Sbjct: | 232 | ITPPKPFIYYDNFKEFEVTICARYTYGKPVPGVAYVRFGVKDEDGKKELLAGLE | 285 |
| Query: | 292 | NLSGQTDKTGCFSAPVDMATFDLIGYAY-SHQINIVATVVEEGTGVEANA-TQNIYIS + | 347 |
| Sbjct: | 286 | ERAKLLDGNGEICLSQEVLLKELQLKNEDLEGKSLYVAVAVIESEGGDMEEAELGGIKIV | 345 |
| Query: | 348 | PQMGSMTFEDTSNFYHPNFPFSGKMLLKFPQGGVLPCKNHLVFLVIYGTNGTFNQTLVTD + | 407 |
| Sbjct: | 346 | RSPYKLKFVKTPSHFKPGIPFFLKVLVVDPDGSPAPNVPVKVSAQDASYYSNGTTD | 401 |
| Query: | 408 | NNGLAPFTLETSGWNGTDVSLEGKFQMEDLVYNPEQVPRYYQNAYLHLRPFYSTTRSFLG + + + + + + + + + + + | 467 |
| Sbjct: | 402 | EDGLAQFSINTSGISSLSITVRTNHKELPEEVQAHAEAQATAYSTVSLSKSYIHLS | 457 |
| Query: | 468 | IHRLNGPLKCGQPQEVLVDYYIDPADASPDQEISFSYYLIGKGSLVMEGQKHLNSKKKGL | 527 |
| Sbjct: | 458 | IERTLPCGPGVGEQANFILRGKSLGELKILHFYYLIMSKGKIVKTGREPREPG | 510 |
| Query: | 528 | KASFSLSLTFTSRLAPDPSLVIYAIFPSGGVVADKIQFSVEMCFDNQQL + + + + | 576 |
| Sbjct: | 511 | QGLFSLSIPVTPDLAPSFRLVAYYILPQGEVVADSVWIDVEDCCANKLDLSFSPSKDYRL | 570 |
| Query: | 577 | PGAEVELQLQAAPGSLCALRAVDESVLLLRPDRELSNRSVY 617 (of SEQ ID NO + + +++ + | :10) |
| Sbjct: | 571 | PAQQVKLRVEADPQSLVALRAVDQAVYLLKPKAKLSMSKVY 611 (SEQ ID NO:81 | <u>)</u> |

At page 52 please replace Table 3E with the following:

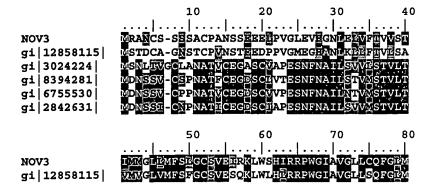
| | Table 3E. | BLAST re | sults for NO | V3 | |
|--|--|-------------|------------------|------------------|--------|
| Gene Index/ Identifier | Protein/ Organism | Length (aa) | Identity (%) | Positives (%) | Expect |
| gi 12858115 dbj BAB 31203.1 (AK018423) BAB31203 | putative [Mus musculus] | 373 | 187/310 (60%) | 226/310 (72%) | 6e-87 |
| gi 3024224 sp Q2872 7 NTCI RABIT NP 058918 | ILEAL SODIUM/BILE ACID COTRANSPORTER (ILEAL NA(+)/BILE ACID COTRANSPORTER) (NA+ DEPENDENT ILEAL BILE ACID TRANSPORTER) (ILEAL SODIUM- DEPENDENT BILE ACID TRANSPORTER) (ISBT) (SODIUM/TAURO CHOLATE COTRANSPORTIN G POLYPEPTIDE, ILEAL) | 347 | 116/279 (41%) | 173/279 (61%) | 1e-52 |
| gi 8394281 ref NP 0 58918.1 NP 035518 | solute carrier family 10, member 2 [Rattus norvegicus] | 348 | 130/344 (37%) | 195/344 (55%) | 2e-52 |
| gi 6755530 ref NP_0 35518.1 NP 35518 | solute carrier family 10, member 2 [Mus musculus] | 348 | 125/313 (39%) | 191/313 (60%) | 4e-52 |
| gi 6755530 ref NP_0 35518.1 NP 35518 | solute carrier family 10, member 2 [Mus musculus] | 348 | 125/313 (39%) | 191/313 (60%) | 4e-52 |

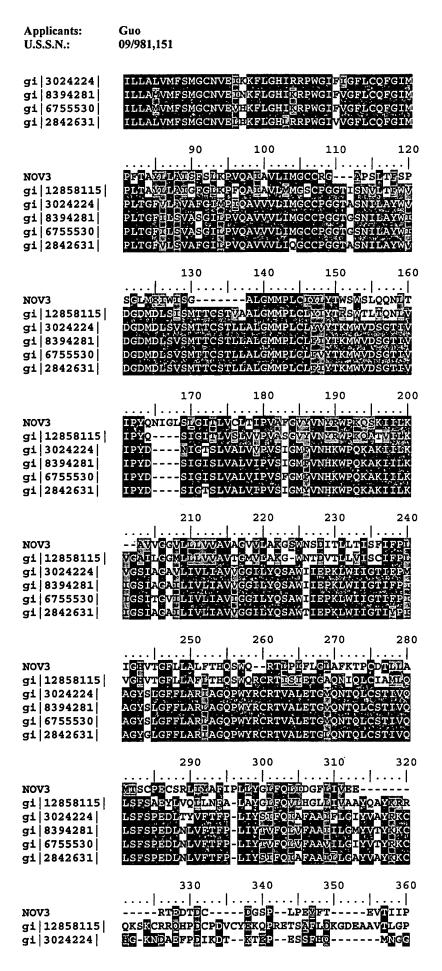
| gi 2842631 sp Q6041 | ILEAL | 348 | 121/306 | 185/306 | 4e-52 |
|---------------------|---------------|-----|---------|---------|-------|
| 4 NTCI CRIGR | SODIUM/BILE | | (39%) | (59%) | |
| Q60414 | ACID | | | | |
| | COTRANSPORTER | | | | 1 |
| | (ILEAL | | | | 1 |
| | NA(+)/BILE | | | | |
| | ACID | | | | [|
| | COTRANSPORTER | | | | |
| |) (NA+ | | | | - |
| | DEPENDENT | | | | |
| | ILEAL BILE | | | | |
| | ACID | | | | |
| | TRANSPORTER) | | | | l |
| | (ILEAL | | | | |
| | SODIUM- | | | | |
| | DEPENDENT | | | | |
| | BILE ACID | | | | |
| | TRANSPORTER) | | | | |
| | (ISBT) | | | | |
| | (SODIUM/TAURO | | | | |
| | CHOLATE | | | · | |
| | COTRANSPORTIN | | | | |
| | G | | | | |
| | POLYPEPTIDE, | | | | |
| | ILEAL) | | | | L |

At page 53 please replace Table 3F with the following:

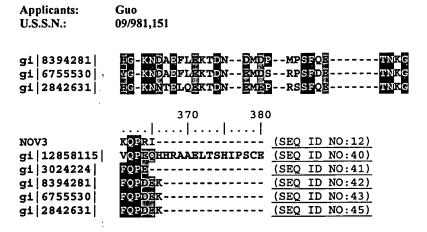
Table 3F. ClustalW Analysis of NOV3

- 1) NOV3 (SEQ ID NO:12)
- 3) $\frac{gi|12858115|dbj|BAB31203.1|}{BAB31203}$ $\frac{gi|12858115|}{BAB31203}$ (AK018423) putative [Mus musculus] (SEQ ID NO:40)
- 4) gi|3024224|sp|Q28727|NTCI RABIT gi|3024224| NP058918 ILEAL SODIUM/BILE ACID COTRANSPORTER (ILEAL NA(+)/BILE ACID COTRANSPORTER) (NA+ DEPENDENT ILEAL BILE ACID TRANSPORTER) (ILEAL SODIUM-DEPENDENT BILE ACID TRANSPORTER) (ISBT) (SODIUM/TAUROCHOLATE COTRANSPORTING POLYPEPTIDE, ILEAL) (SEQ ID NO:41)
- 5) <u>gi|8394281|ref|NP 058918.1|</u> <u>gi|8394281| NP035518</u> solute carrier family 10, member 2 [Rattus norvegicus] (SEQ ID NO:42)
- 6) <u>gi|6755530|ref|NP 035518.1|</u> <u>gi|6755530| NP035518</u> solute carrier family 10, member 2 [Mus musculus] (SEQ ID NO:43)
- 7) $\frac{gi|6755530|ref|NP|035518.1|}{member 2 [Mus musculus]} \frac{gi|6755530|}{(SEQ ID NO:44)}$ NP035518 solute carrier family 10,
- 8) gi|2842631|sp|Q60414|NTCI_CRIGR_g1|2842631| Q60414 ILEAL SODIUM/BILE ACID COTRANSPORTER (ILEAL NA(+)/BILE ACID COTRANSPORTER) (NA+ DEPENDENT ILEAL BILE ACID TRANSPORTER) (ILEAL SODIUM-DEPENDENT BILE ACID TRANSPORTER) (ISBT) (SODIUM/TAUROCHOLATE COTRANSPORTING POLYPEPTIDE, ILEAL) (SEQ ID NO:45)









Page 55, please replace Table 3G with the following

Table 3G Domain Analysis of NOV3

gnl|Pfam|pfam01758, SBF, Sodium Bile acid symporter family. This
family consists of Na+/bile acid co-transporters. These transmembrane
proteins function in the liver in the uptake of bile acids from portal
blood plasma a process mediated by the co-transport of Na+. Also in
the family is ARC3 from S. cerevisiae this is a putative transmembrane
protein involved in resistance to arsenic compounds. (SEQ ID NO:82)
CD-Length = 186 residues, 80.1% aligned
Score = 75.9 bits (185), Expect = 3e-15

```
TIMMGLLMFSLGCSVEIRKLWSHIRRPWGIAVGLLCQFGLMPFTAYLLAIS-FSLKPVQA
Query:
       39
                                   +||| + +||| |+ +||
             + + |+|||+|
                              - 1
                                                       ++||
            ALGLFLMMFSMGLKVRFEDLKEALRRPKALILGLLLQWIIMPLLMFILAWLLLRLPPELA
Sbjct:
       1
            IAVLIMGCCRG---APSLTFSPSGLMEIWIS-----GALGMMPLCIYLYTWSWSLQQN
Query:
       98
                              |+
                                   | +|+ +
                                                     + || +|
            TGLILVGCAPGGAMSNVWTYLAKGDVELSVVMVALSTLLAPLVTPLLSFLLAGLL-
Sbjct:
            LTIPYQNIGLSLGITLVCLTIPVAFGVYVNYRWP 181 (of SEQ ID NO:12)
Query:
                       + || + ||+ |+
Sbjct:
       116 VHVDAVSPWSLIKSVLVYVIIPLIAGMLTRYFLP
                                                     (SEQ ID NO 82)
                                                149
```

At page 61, please replace Table 4C with the following:

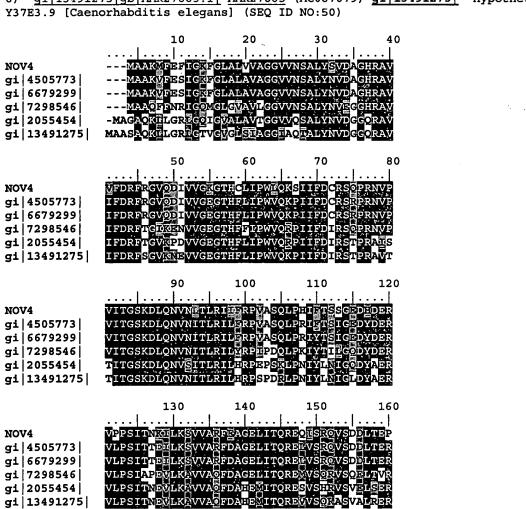
| | Table 4C. BLA | ST result | ts for NOV | 1 | |
|---|---------------------------|-------------|------------------|------------------|--------|
| Gene Index/ Identifier | Protein/ Organism | Length (aa) | Identity (%) | Positives (%) | Expect |
| gi 4505773 ref NP 0 02625.1 NP 002625 | prohibitin [Homo sapiens] | 272 | 236/270 (87%) | 251/270 (92%) | e-123 |
| gi 92643 pir A3968 2 A39682 | prohibitin - rat | 272 | 235/270 (87%) | 251/270 (92%) | e-123 |

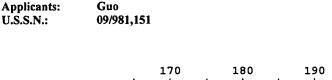
| gi 7298546 gb AAF53 765.1 (AE003661) AAF53765 | 1(2)37Cc gene product [Drosophila melanogaster] | 276 | 178/270 (65%) | 220/270 (80%) | 8e-96 |
|---|--|-----|------------------|------------------|-------|
| gi 2055454 gb AAB53 231.1 (U97204) AAB53231 | prohibitin-like molecule TC-PRO-1 [Toxocara canis] | 274 | 153/264 (57%) | 209/264 (78%) | 9e-80 |
| gi 13491275 gb AAK2 7865.1 (AC087079) AAK27865 | Hypothetical protein Y37E3.9 [Caenorhabditis elegans] | 275 | 154/270 (57%) | 210/270 (77%) | 6e-79 |

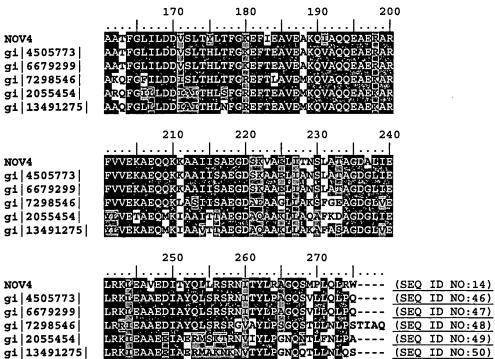
At page 61 please replace Table 4D with the following:

Table 4D ClustalW Analysis of NOV4

- NOV4 (SEQ ID NO:14) gi|4505773|ref|NP_002625.1| NP_002625 gi|4505773| prohibitin [Homo sapiens] (SEQ 2) ID NO:46)
- prohibitin rat (SEQ ID NO:47) 3) gi|92643|pir||A39682 A39682 gi|6679299|
- gi|7298546|gb|AAF53765.1| AAF53765 (AE003661) gi|7298546| 1(2)37Cc gene product [Drosophila melanogaster] (SEQ ID NO:48)
- 5) gi|2055454|gb|AAB53231.1| AAB53231 (U97204) gi|2055454| prohibitin-like molecule TC-PRO-1 [Toxocara canis] (SEQ ID NO:49)
- 6) <u>gi|13491275|gb|AAK27865.1|</u> AAK27865 (AC087079) <u>gi|13491275|</u> Hypothetical protein







At page 62 please replace Table 4E with the following:

Table 4E. Domain Analysis of NOV4

gnl|Pfam()pfam()1145, Band_7, SPFH domain / Band 7 family. This family
also includes proteins with high blast scores to known Band 7 protein:
HflC from E. coli HflK from E. coli, and Prohibitin family members
(SEQ ID NO:83)
CD-Length = 191 residues, 91.6% aligned

CD-Length = 191 residues, 91.6% aligned Score = 157 bits (397), Expect = 7e-40

| Query: | 28 | YSVDAGHRAVVFDRFRGVQDIVVGKGTHCLIPWLQKSIIFDCRSQPRNVPVITGSKDLQN | 87 |
|--------|-------|--|-----|
| Sbjct: | 17 | KIVKEYERGVIFRLGRYVRQ-VVGPGLHFIIPFIDTVKKVDLRTVVYDVPSQEIITKDNV | 75 |
| Query: | 88 | VNLTLRIIFRPVASQLPHIFTSSGEDHDERVPPSITNKILKSVVARFEAGELITQREQIS | 147 |
| Sbjct: | 76 | | 132 |
| Query: | 148 | RQVSDDLTEPAATFGLILDDVSLTYLTFGKEFIEAVEAKQIAQQEAERARFVVEKAEQQK | 207 |
| Sbjct: | 133 | SQLREILDEATDPWGIKVERVEIKDIRLPEEVQRAMAAQMEAEREA-RAKILEAEGEQEA | 191 |
| Query: | (aa 2 | 8-207 of SEQ ID NO:14) | |

Sbjct: (SEQ ID NO:83)

At page 63, please replace Table 4F with the following:

Table 4F. Domain Analysis of NOV4

gnl|Smart|smart00244, PHB, prohibitin homologues; prohibitin
homologues (SEQ ID NO:84)
CD-Length = 160 residues, 98.8% aligned
Score = 97.4 bits (241), Expect = 9e-22

YSVDAGHRAVVFDRFRGVQDIVVGKGTHCLIPWLQKSIIFDCRSQPRNVPVITG-SKDLQ Query: 28 | |+| | +||++ | +| | |+| +|| FYVIGEGERGVVERLGRVLK-VLGPGLHFVIPFIDDVKRVDLRAQTDDVPPQEVITKDNV Sbjct: Query: NVNLTLRIIFRPVASQLPHIFTSSGEDHDERVPPSITNKILKSVVARFEAGELIT-QREQ ... |+||+ + TVSVDAVVYYR-VLDPLKAVYGV--LDADYRALROLAQTTLRSVIGKRTLDELLTDEREK 118 Sbjct: 62 ISRQVSDDLTEPAATFGLILDDVSLTYLTFGKEFIEAVEAKQ 187 (of SEQ ID NO:14) Query: 146 || + ++| | | +|+ ++|| + + +| ||+||+| Sbjct: 119 ISENIREELNEAAEPWGIEVEDVEIKDIRLPEEIKEAMEAQQ 160 (SEQ ID NO:84)

At page 67 please replace Table 5C with the following:

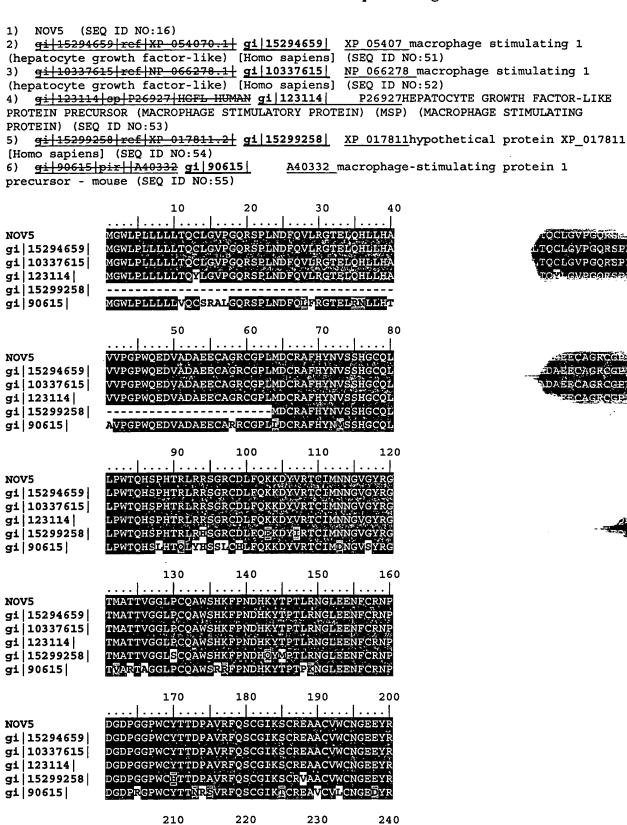
| Table 5C. BLAST results for NOV5 | | | | | |
|---|---|-------------|------------------|------------------|--------|
| Gene Index/ Identifier | Protein/ Organism | Length (aa) | Identity (%) | Positives (%) | Expect |
| 91 15294659 ref XP 054070.1 XP 05407 | macrophage stimulating 1 (hepatocyte growth factor- like) [Homo sapiens] | 711 | 561/720 (77%) | 577/720 (79%) | 0.0 |
| gi 10337615 ref NP 066278.1 NP 066278 | macrophage stimulating 1 (hepatocyte growth factor- like) [Homo sapiens] | 711 | 560/720 (77%) | 576/720 (79%) | 0.0 |
| gi 123114 sp P26927 HGFL_HUMAN P26927 | HEPATOCYTE GROWTH FACTOR-LIKE PROTEIN PRECURSOR (MACROPHAGE STIMULATORY PROTEIN) (MSP) (MACROPHAGE STIMULATING PROTEIN) | 711 | 560/720 (77%) | 576/720 (79%) | 0.0 |
| gi 15299258 ref XP 017811.2 XP 017811 | hypothetical protein XP_017811 [Homo sapiens] | 529 | 440/532 (82%) | 456/532 (85%) | 0.0 |
| gi 90615 pir A4033 2 A40332 | macrophage- stimulating protein 1 precursor - mouse | 716 | 449/725 (61%) | 507/725 (69%) | 0.0 |

Applicants: U.S.S.N.:

Guo 09/981,151

At page 68 please replace Table 5D with the following:

Table 5D Clustal W Sequence Alignment



DAEECAGREGELMOCRAFEE

HANGER CHEST

DAEECAGRCGELMEGE?

TOCLGYPGORSPLNDFOWLRGFELC

OCHGVPGORS



| U.S.S.N.: | 09/981,151 |
|--|--|
| NOV5 gi 15294659 gi 10337615 gi 123114 gi 15299258 gi 90615 | GAVDRTESGRECORWDLOHPHOHPFEPGKFLDOGLDDNYC GAVDRTESGRECORWDLOHPHOHPFEPGKFLDOGLDDNYC GAVDRTESGRECORWDLOHPHOHPFEPGKFLDOGLDDNYC GAVDRTESGRECORWDLOHPHOHPFEPGKFLDOGLDDNYC GAVDRTESGRECORWDLOHPHOHPFEPGKFLDOGLDDNYC GAVDRTESGRECORWDLOHPHOHPFEPGKFLDOGLDDNYC GEVDVTESGRECORWDLOHPHSHPFEPGKFLDKDLKDNYC |
| | 250 260 270 280 |
| NOV5 gi 15294659 gi 10337615 gi 123114 gi 15299258 gi 90615 | RNPDGSERPWCYTTDPOIEREFCDLPRCG-SE RNPDGSERPWCYTTDPOIEREFCDLPRCG-SE RNPDGSERPWCYTTDPOIEREFCDLPRCG-SE RNPDGSERPWCYTTDPOIEREFCDLPRCG-SE RNPDGSERPWCYTTDPOIEREFCDLPRCG-SE RNPDGSERPWCYTTDPOIEREFCDLPRCG-SE RNPDGSERPWCYTTDPOIEREFCDLPRCG-SE |
| | 290 300 310 320 |
| NOV5 gi 15294659 gi 10337615 gi 123114 gi 15299258 gi 90615 | AOPROEATTVSCFRGKGEGYRGTANTTTAGVPCORWDAOI AOPROEATTVSCFRGKGEGYRGTANTTTAGVPCORWDAOI AOPROEATTVSCFRGKGEGYRGTANTTTAGVPCORWDAOI AOPROEATTVSCFRGKGEGYRGTANTTTAGVPCORWDAOI AOPROEATTVSCFRGKGEGYRGTANTTTAGVPCORWDAOI AOPROEATTVSCFRGKGEGYRGTANTTTAGVPCORWDAOI SORRMAGKANNCFRGKGEDYRGT |
| | 330 340 350 360 |
| NOV5 gi 15294659 gi 10337615 gi 123114 gi 15299258 gi 90615 | PHOHRFTPEKYACKOLRENFCRNPDGSEAPWCFTLRPGMR PHOHRFTPEKYACKOLRENFCRNPDGSEAPWCFTLRPGMR PHOHRFTPEKYACKOLRENFCRNPDGSEAPWCFTLRPGMR PHOHRFTPEKYACKOLRENFCRNPDGSEAPWCFTLRPGMR PHOHRFTPEKYACKOLRENFCRNPDGSEAPWCFTLRPGMR PHOHRFTPEKYACKOLRENFGRNPDGSEAPWCFTLRPGMR |
| | 370 380 390 400 |
| NOV5 gi 15294659 gi 10337615 gi 123114 gi 15299258 gi 90615 | AAFCYOIRRCTDDVREOTATTAOGSSTAARSARPARVSSA AAFCYOIRRCTDDVREODCYHGAGEOYRGTVSKTRKGVOC AAFCYOIRRCTDDVREODCYHGAGEOYRGTVSKTRKGVOC AAFCYOIRRCTDDVREODCYHGAGEOYRGTVSKTRKGVOC VGFCYOIRRCTDDVREODCYHGAGEOYRGTVSKTRKGVOC MAFCEOIPRCTETTVPEGCYHGEGEOYRGTVSKTRKGVOC |
| | 410 420 430 440 |
| NOV5 gi 15294659 gi 10337615 gi 123114 gi 15299258 gi 90615 | SAGPLRRRTSRSSRLPPNRMENWRRTSAGTOMG I AMGEGA QRWSAETPHKPOFTFTSEPHAQLEENFCRNPDGDSHGP QRWSAETPHKPOFTFTSEPHAQLEENFCRNPDGDSHGP QRWSAETPHKPOFTFTSEPHAQLEENFCRNPDGDSHGP QRGSAETPHKPOFTFTSEPHAQLEENFCRNPDGDSHGP |
| NOV5 gi 15294659 gi 10337615 gi 123114 gi 15299258 gi 90615 | 450 460 470 480 TRNTOGPHSTTVPCDAALMTSRHQSWTPQTRCSLRSVA WCYTTOPRIPPDYCALRCADDQPPSILDPPDQVQFEKCG WCYTTOPRIPPDYCALRCADDQPPSILDPPDQVQFEKCG WCYTTOPRIPPDYCALRCADDQPPSILDPPDQVQFEKCG TRNTQGPHSTTVPCDAALMTSRHQSWTPQTRCSLRSVA WCYTTPDPDILFDYCALPRCDDDQPPSILDPPDQVVFEKCG |
| | 490 500 510 520 |

Applicants:

U.S.S.N.: 09/981,151 RGWIGWISGVPSCAWEGAIRATHEGQSACGIG-----NOV5 KRVDRLDORRSKLRVVGGHPGNSPWTVSLRNROGOHFCGG KRVDRLDORRSKLRVVGGHPGNSPWTVSLRNROGOHFCGG KRVDRLDORRSKLRVVGGHPGNSPWTVSLRNROGOHECGG KRVDRLDORRSKLRVVGGHPGNSPWTVSLRNROGOHECGG K----RVDRSNKLRVVGGHPGNSPWTVSLRNROGOHFCGG gi | 15294659 | gi | 10337615 | gi | 123114 | gi | 15299258 | gi|90615| 530 540 550 -----MEPLTGYEVWLGTLFQNPQHGE NOV5 MEPLICYEVWLGTLFONPONGE
SLVKEOWILTAROCFSSCHMPLTGYEVWLGTLFONPONGE
SLVKEOWILTAROCFSSCHMPLTGYEVWLGTLFONPONGE
SLVKEOWILTAROCFSSCHMPLTGYEVWLGTLFONPONGE
SLVKEOWILTAROCFSSCHMPLTGYEVWLGTLFONPONGE
SLVKEOWILTAROCFSSCHMPLTGYEVWLGTLFONPONGE
SLVKEOWILTAROCTWSCHEPLTGYEVWLGTLFONPONGE gi | 15294659 | gi | 10337615 | gi | 123114 | gi|15299258| gi | 90615 | 580 590 . | | | | | | | . PSLORVPVAKMYCGPSGSQLVLLKLERSVTLNORVALICL
PSLORVPVAKMYCGPSGSQLVLLKLERSVTLNORVALICL
PSLORVPVAKMYCGPSGSQLVLLKLERSVTLNORVALICL
PSLORVPVAKMYCGPSGSQLVLLKLERSVTLNORVALICL
PGLORVPVAKMECGPSGSQLVLLKLERSVTLNORVALICL
ATLORVPVAKMOCGPEGSQLVLLKLEREVTLNORVALICL NOV5 gi | 15294659 | gi | 10337615 | gi | 123114 | g1|15299258| gi | 90615 | PPEWYVVPPGTKCEIAGWGETKGTGNDTVLNVALLNVISN NOV5 PPEWYVVPPGTKCEIAGWGETKGTGNDTVLNVA<mark>L</mark>LNVISN gi | 15294659 | PPEWYVVPPGTKCEIAGWGETKGTGNDTVLNVA<mark>F</mark>LNVISN PPEWYVVPPGTKCEIAGWGETKGTGNDTVLNVA<mark>F</mark>LNVISN gi | 10337615 | gi | 123114 | gi | 15299258 | PPEOYVVPPGTKCEIAGWGESIGTSNYTVLEVASYNVISN gi | 90615 | 660 NOV5 gi | 15294659 | QECNIKHRGRVRESENCTEGI APVGACEGDYGGPI QECNIKHRGRVRESENCTEGI APVGACEGDYGGPI QECNIKHRGRVRESENCTEGI APVGACEGDYGGPI gi | 10337615 | gi | 123114 | gi | 15299258 | QECN<mark>TKYRGER©ESEECT©GLYV</mark>PVGACEGDYGGPLACYT gi | 90615 | 700 710 720 . | | | . . | HNCWVLEGIIIPNRVCARS<mark>C</mark>WPAVFTRVSVFVDWIHKVMR NOV5 HNCWVLEGIIIPNRVCARSRWPAVFTRVSVFVDWIHKVMR qi | 15294659 | HNCWVLEGITÎ PNRVCAŘSRWPAVFTRVSVFVDWIHKVMŘ gi | 10337615 | HNCWVLEGIIIPNRVCARSRWPAVFTRVSVFVDWIHKVMF gi | 123114 | gi | 15299258 | _____ hecwvlegeiipnrvcar<mark>p</mark>rwpaeftrvsvfvdwi<u>kv</u>we gi | 90615 | NOV5 (SEQ ID NO:16) gi | 15294659 | (SEQ ID NO:51) (SEQ ID NO:52) gi | 10337615 | gi | 123114 | (SEQ ID NO:53) gi | 15299258 | (SEQ ID NO:54) (SEQ ID NO:55) gi|90615|

Applicants:

Guo

GWGETKOTONTVLAWZ AGWGETKGTGNDTVLAWZ EIAGWGETKGTGNDTVLAVZ

At pages 70, please replace Table 5E with the following:

Table 5E. Domain Analysis of NOV5

gnl|Pfam|pfam00051, kringle, Kringle domain. Kringle domains have been
found in plasminogen, hepatocyte growth factors, prothrombin, and
apolipoprotein A. Structure is disulfide-rich, nearly all-beta. (SEQ
ID NO:85)

CD-Length = 79 residues, 100.0% aligned Score = 117 bits (292), Expect = 3e-27

Query: 250 WCYTTDPQIEREFCDLPRC 268 (of SEQ ID NO:16)

Sbjct: 61 WCYTTDPRVRWEYCDIPRC 79 (SEQ ID NO:85)

At pages 71, please replace Table 5F, with the following:

Table 5F. Domain Analysis of NOV5

gnl | Pfam | pfam00051, kringle, Kringle domain. Kringle domains have been found in plasminogen, hepatocyte growth factors, prothrombin, and apolipoprotein A. Structure is disulfide-rich, nearly all-beta. (SEQ ID NO:85)

CD-Length = 79 residues, 100.0% aligned Score = 112 bits (279), Expect = 9e-26

Query: 342 WCFTLRPGMRAAFCYQIRRC 361 (of SEQ ID NO:16)

Sbjct: 61 WCYTTDPRVRWEYC-DIPRC 79 (SEQ ID NO:85)

At pages 71, please replace Table 5G with the following:

Applicants: U.S.S.N.:

Guo 09/981,151

Table 5G. Domain Analysis of NOV5

gnl|Pfam|pfam00051, kringle, Kringle domain. Kringle domains have been found in plasminogen, hepatocyte growth factors, prothrombin, and apolipoprotein A. Structure is disulfide-rich, nearly all-beta. (SEQ ID NO:85)

CD-Length = 79 residues, 100.0% aligned Score = 104 bits (259), Expect = 2e-23

Sbjct: 1 CYHGNGENYRGTASTTESGAPCQRWDSQTPHRHSKYTPERYPAKGLGENYCRNPDGDE-R 59

Query: 167 PWCYTTDPAVRFQSCGIKSC 186 (of SEQ ID NO:16)

At pages 71-72, please replace Table 5H with the following:

Table 5H. Domain Analysis of NOV5

gnl|Smart|smart00130, KR, Kringle domain; Named after a Danish pastry.
Found in several serine proteases and in ROR-like receptors. Can occur
in up to 38 copies (in apolipoprotein(a)). Plasminogen-like kringles
possess affinity for free lysine and lysine- containing peptides.
(SEQ ID NO:86)

CD-Length = 83 residues, 97.6% aligned Score = 111 bits (278), Expect = 1e-25

Query: 191 CVWCNGEEYRGAVDRTESGRECQRWDLQHPHQHPFEPGKFLDQGLDDNYCRNPDG-SERP 249

Sbjct: 3 CYAGNGESYRGTASTTKSGKPCQRWDSQTPHLHRFTPERFPELGLEHNYCRNPDGDSEGP 62

Query: 250 WCYTTDPQIEREFCDLPRCGS 270 (of SEQ ID NO:16)

Sbjct: 63 WCYTTDPNVRWEYCDIPQCES 83 (SEQ ID NO:)

At pages 72, please replace Table 5I with the following:

Applicants: U.S.S.N.:

Guo 09/981,151

Table 5I. Domain Analysis of NOV5

gnl|Smart|smart00130, KR, Kringle domain; Named after a Danish pastry.
Found in several serine proteases and in ROR-like receptors. Can occur
in up to 38 copies (in apolipoprotein(a)). Plasminogen-like kringles
possess affinity for free lysine and lysine- containing peptides.
(SEO ID NO:86)

CD-Length = 83 residues, 97.6% aligned Score = 106 bits (265), Expect = 4e-24

Query: 108 RTCIMNNGVGYRGTMATTVGGLPCQAWSHKFPNDHKYTPTLRN--GLEENFCRNPDGDPG 165

Sbjct: 1 RDCYAGNGESYRGTASTTKSGKPCQRWDSQTPHLHRFTPERFPELGLEHNYCRNPDGDSE 60

Query: 166 GPWCYTTDPAVRFQSCGIKSC 186 (of SEQ ID NO:16)

At pages 72, please replace Table 5J with the following:

Table 5J. Domain Analysis of NOV5

gnl|Smart|smart00130, KR, Kringle domain; Named after a Danish pastry.
Found in several serine proteases and in ROR-like receptors. Can occur
in up to 38 copies (in apolipoprotein(a)). Plasminogen-like kringles
possess affinity for free lysine and lysine- containing peptides.
(SEQ ID NO:86)

CD-Length = 83 residues, 97.6% aligned Score = 104 bits (260), Expect = 1e-23

Query: 283 CFRGKGEGYRGTANTTTAGVPCQRWDAQIPHQHRFTPEKYACKDLRENFCRNPDG-SEAP 341

Sbjct: 3 CYAGNGESYRGTASTTKSGKPCQRWDSQTPHLHRFTPERFPELGLEHNYCRNPDGDSEGP 62

Query: 342 WCFTLRPGMRAAFCYQIRRCTD 363 (of SEQ ID NO:16)

Sbjct: 63 WCYTTDPNVRWEYCD-IPQCES 83 (SEQ ID NO:86)

||+| | +| +| | +|

At pages 72-73, please replace Table 5K with the following:

Table 5K. Domain Analysis of NOV5

gnl|Smart|smart00020, Tryp_SPc, Trypsin-like serine protease; Many of
these are synthesised as inactive precursor zymogens that are cleaved
during limited proteolysis to generate their active forms. A few,
however, are active as single chain molecules, and others are inactive
due to substitutions of the catalytic triad residues. (SEQ ID NO:87)
CD-Length = 230 residues, 79.1% aligned
Score = 110 bits (274), Expect = 3e-25

```
504 PLTGYEVWLGTLFQNPQHGEPSLQRVPVAKMVCGPSGSQ-----LVLLKLERSVTLNQR 557
Query:
                 | | | + +
                                  | | |+|++ |+ +
                                                       + | | | | |
                                                               |||+
            APSSIRVRLGSHDLSSGEET - - - QTVKVSKVIVHPNYNPSTYDNDIALLKLSEPVTLSDT
Sbjct:
       49
            VALICLPPEWYVVPPGTKCEIAGWGETKGTG--NDTVLNVALLNVISNQECNIKHRGR--
Query:
       558
            | | | | | | | | | | ++||| | + | | + ++|| | | + |
            VRPICLPSSGYNVPAGTTCTVSGWGRTSESSGSLPDTLQEVNVPIVSNATCRRAYSGGPA
Sbjct:
       106
                                                                        165
            -----GDYGGPLACFTHNCWVLEGIIIPNRV-CARSCWPAVFTRVS
Query:
       614
                               ||| ||+
                                                         | |+|||
            ITDNMLCAGGLEGGKDACQGDSGGPLVC-NDPRWVLVGIVSWGSYGCARPNKPGVYTRVS
Sbjct:
       166
            VFVDWI
                   659
                        (of SEQ ID NO:16)
Query:
       654
             ++|||
                   230
Sbjct:
       225
            SYLDWI
                         (SEQ ID NO:87)
```

At pages 73, please replace 5L with the following:

Table 5L. Domain Analysis of NOV5

gnl|Pfam|pfam00089, trypsin, Trypsin. Proteins recognized include all
proteins in families S1, S2A, S2B, S2C, and S5 in the classification
of peptidases. Also included are proteins that are clearly members,
but that lack peptidase activity, such as haptoglobin and protein Z
(PRTZ*). (SEQ ID NO:88)
CD-Length = 217 residues, 77.9% aligned
Score = 92.0 bits (227), Expect = 9e-20

| Query: | 508 | YEVWLGTLFQNPQHGEPSLQRVPVAKMVCGPSGSQLVLLKLERSVTLNQRVALICL | 563 |
|----------|-------|--|-----|
| Obd === | 4.0 | | 106 |
| SDJCt: | 49 | VRVVLGEHNLGTTEGTEQKFDVKKIIVHPNYNPDTNDIALLKLKSPVTLGDTVRPICL | 106 |
| Query: | 564 | PPEWYVVPPGTKCEIAGWGETKGTGNDTVLNVALLNVISNQECNIKHRG | 612 |
| Sbjct: | 107 | PSASSDLPVGTTCSVSGWGRTKNLGTSDTLQEVVVPIVSRETCRSAYGGTVTDTMICAGA | 166 |
| Query: | 613 | RGDYGGPLACFTHNCWVLEGIIIPNRVCARSCWPAVFTRVSVFVDWI 659 + | |
| Sbjct: | 167 | LGGKDACQGDSGGPLVCSDGELVGIVSWGYGCAVGNYPGVYTRVSRYLDWI 217 | |
| 01102771 | lof c | FO ID NO.16) | |

Query: (of SEQ ID NO:16)
Sbjct: (SEQ ID NO:88)

At pages 73, please replace Table 5M with the following:

Table 5M. Domain Analysis of NOV5

gnl|Smart|smart00473, PAN_AP, divergent subfamily of APPLE domains; Apple-like domains present in Plasminogen, C. elegans hypothetical ORFs and the extracellular portion of plant receptor-like protein kinases. Predicted to possess protein- and/or carbohydrate-binding functions. (SEQ ID NO:89) CD-Length = 79 residues, 94.9% aligned Score = 52.0 bits (123), Expect = 1e-07

Query: DFQVLRGTELQHLLHAVVPGPWQEDVADAEECAGRC-GPLMDCRAFHYNVSSHGCQLLPW | | |+| || |||| +| ||+| || + || | CFVRLPNTKL-----PDFSPIVISVASLEECAQKCLNSNCSCRSFTYNNDTKGCLLWSE 5 Sbjct: TQHSPHTRLRRSGRCDLFQKK 104 (of SEQ ID NO:16) Query: 84 Sbjct: 59 SSLGDARQLLPSGGVDYYEKI 79 (SEQ ID NO:89)

At pages 74, please replace Table 5N with the following:

Table 5N. Domain Analysis of NOV5

gnl|Pfam|pfam00024, PAN, PAN domain. The PAN domain contains a
conserved core of three disulphide bridges. In some members of the
family there is an additional fourth disulphide bridge the links the N
and C termini of the domain. The domain is found in diverse proteins,
in some they mediate protein-protein interactions, in others they
mediate protein-carbohydrate interactions. (SEQ ID NO:159)
CD-Length = 78 residues, 76.9% aligned
Score = 50.1 bits (118), Expect = 4e-07

| Query: | 45 | PWQEDVADAEECAGRCGPLMD-CRAFHYNVSSHGCQLLPWTQHSPHTRLRRSGRCDLFQK | 103 |
|--------|------|--|-----|
| Sbjct: | 17 | + + + + ++ ISVVNVPSLEECAALCLEEPRVCRSFTYNNKSKQCLLKSESSGSLPRLKRPSQKVDYYEK | |
| Query: | (of | SEQ ID NO:16) | |
| Sbjct: | (SEC | Q ID NO:159) | |

At page 78 please replace Table 6C with the following:

Applicants: U.S.S.N.:

Guo 09/981,151

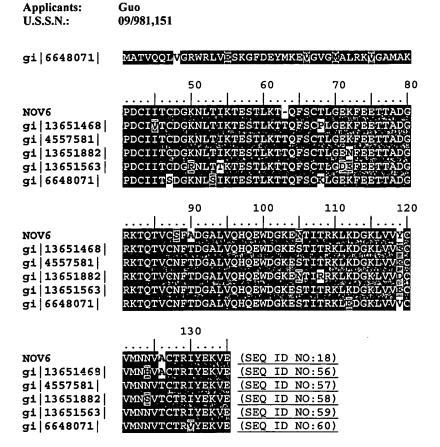
| Table 6C. BLAST results for NOV6 | | | | | |
|--|---|----------------|------------------|------------------|--------|
| Gene Index/ Identifier | Protein/ Organism | Length (aa) | Identity (%) | Positives (%) | Expect |
| gi 13651468 ref XP 016351.1 XP 016351 | similar to GASTRIN/CHOLECYST OKININ TYPE B RECEPTOR (CCK-B RECEPTOR) (CCK-BR) (H. sapiens) [Homo sapiens] | 135 | 122/135 (90%) | 126/135 (92%) | 1e-58 |
| gi 4557581 ref NP 0 01435.1 NP 001435 | fatty acid binding protein 5 (psoriasis- associated); E- FABP [Homo sapiens] | 135 | 124/135 (91%) | 126/135 (92%) | 1e-58 |
| gi 13651882 ref XP 011655.5 XP 011655 | fatty acid binding protein 5 (psoriasis- associated) [Homo sapiens] | 135 | 119/135 (88%) | 124/135 (91%) | 6e-57 |
| gi 13651563 ref XP 015760.1 XP 015760 | similar to GASTRIN/CHOLECYST OKININ TYPE B RECEPTOR (CCK-B RECEPTOR) (CCK-BR) (H. sapiens) [Homo sapiens] | 135 | 118/135 (87%) | 125/135 (92%) | 5e-56 |
| gi 6648071 sp P5505 2 FABE_BOVIN P55052 | FATTY ACID- BINDING PROTEIN, EPIDERMAL (E- FABP) | 135 | 117/135 (86%) | 124/135 (91%) | 1e-55 |

At page 79 please replace Table 6D with the following:

Table 6D Information for the ClustalW proteins

- 1) NOV6 (SEQ ID NO:18)
- 2) gi|13651468|ref|XP 016351.1| gi|13651468| XP 016351similar to GASTRIN/CHOLECYSTOKININ TYPE B RECEPTOR (CCK-B RECEPTOR) (CCK-BR) (H. sapiens) [Homo sapiens] (SEQ ID NO:56)
- 3) $\frac{gi|4557581|ref|NP-001435.1|}{gi|4557581|}$ $\frac{gi|4557581|}{gi|4557581|}$ NP 001435 fatty acid binding protein 5 (psoriasis-associated); E-FABP [Homo sapiens] (SEQ ID NO:57)
- 4) gi|13651882|ref|XP 011655.5| gi|13651882| XP 011655 (psoriasis-associated) [Homo sapiens] (SEQ ID NO:58)
- 5) gi|13651563|ref|XP 015760.1| gi|13651563| XP 015760similar to GASTRIN/CHOLECYSTOKININ TYPE B RECEPTOR (CCK-B RECEPTOR) (CCK-BR) (H. sapiens) [Homo sapiens] (SEQ ID NO:59)
- 6) gi|6648071|sp|P55052|FABE BOVIN gi|13651563| P55052 FATTY ACID-BINDING PROTEIN, EPIDERMAL (E-FABP) (SEQ ID NO:60)





Page 80, please replace Table 6E with the following:

Table 6E. Domain Analysis of NOV6

gnl|Pfam|pfam00061, lipocalin, Lipocalin / cytosolic fatty-acid binding protein family. Lipocalins are transporters for small hydrophobic molecules, such as lipids, steroid hormones, bilins, and retinoids. Alignment subsumes both the lipocalin and fatty acid binding protein signatures from PROSITE. This is supported on structural and functional grounds. Structure is an eight-stranded beta barrel. (SEQ ID NO:90) CD-Length = 145 residues, 76.6% aligned Score = 53.1 bits (126), Expect = 1e-08

```
GRWRLVDSKRFDEYMK-EGGVGTALRKMDAMAK-PDCIITCDGKNLTIKTESTLKTOFSC
Query: 9
            |+| || | || +| || ||
                                           | + | ||
                                                          | |+ | + +
            GKWYLVASANFDPELKEELGVLEATRKEITPLKEGNLEIVFDGDKNGICEETFGKLEKTK 63
Sbjct: 4
Query:
            TLGEKFEETTADGRKTQTVCSFADGALVQHQEWDGKENTITRKLKDGKLVV 117
                                + + || |+ || | + | +|
             | | + | + | | |
            KLGVEFDYYTGDNRFVVLDTDYDNYLLVCVQKGDGNETSRTAELYGRTPEL 114
Sbjct:
       (of SEQ ID NO:18)
Query:
Sbjct: (SEQ ID NO:90)
```

Applicants: U.S.S.N.:

Guo 09/981,151

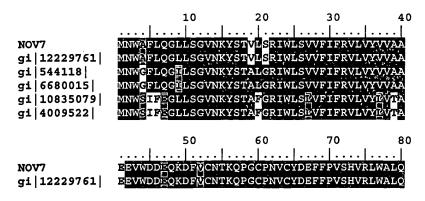
At page 84 please replace Table 7C with the following:

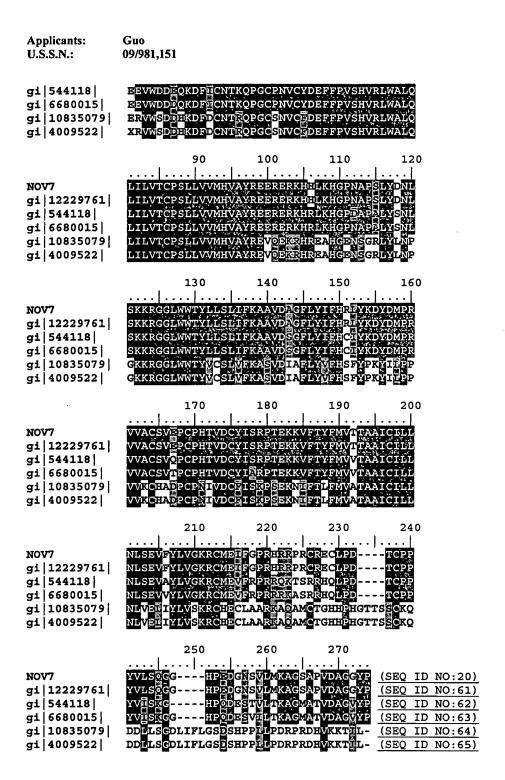
| Table 7C. BLAST results for NOV7 | | | | | | |
|--|--|-------------|-------------------|-------------------|--------|--|
| Gene Index/ Identifier | Protein/ Organism | Length (aa) | Identity (%) | Positives (%) | Expect | |
| 9 12229761 sp Q9NTQ 9 CXB4 HUMAN Q9NTQ9 | GAP JUNCTION BETA-4 PROTEIN (CONNEXIN 30.3) (CX30.3) | 266 | 266/266 (100%) | 266/266 (100%) | e-155 | |
| gi 544118 sp P36380 CXB5 RAT P36380 | GAP JUNCTION BETA-5 PROTEIN (CONNEXIN 30.3) (CX30.3) | 265 | 230/266 (86%) | 245/266 (91%) | e-132 | |
| gi 6600015 ref NP_03 2153.1 NP_032153 | gap junction membrane channel protein beta 4; connexin 30.3 [Mus musculus] | 266 | 230/266 (86%) | 244/266 (91%) | e-131 | |
| gi 10835079 ref NP-0 05259.1 NP 05259 | gap junction protein, beta 5 (connexin 31.1) [Homo sapiens] | 273 | 153/226 (67%) | 177/226 (77%) | 3e-88 | |
| gi 4009522 gb AAC954 72.1 (AF099731)AAC95472 | connexin 31.1 [Homo sapiens] | 273 | 152/226 (67%) | 176/226 (77%) | 1e-87 | |

At page 85 please replace Table 7D with the following:

Table 7D. Information for the ClustalW proteins

- 1) NOV7 (SEQ ID NO:20)
- 2) gi | 12229761 | sp | Q9NTQ9 | CXB4 HUMAN Q9NTQ9 GAP JUNCTION BETA-4 PROTEIN (CONNEXIN 30.3) (CX30.3) (SEQ ID NO:61)
- 3) gi+544118|op+P36380|CXB5-RAT Q9NTQ9 GAP JUNCTION BETA-5 PROTEIN (CONNEXIN 30.3) (CX30.3) (SEQ ID NO:62)
- 4) gi|6680015|ref|NP-032153.1| NP 032153gap junction membrane channel protein beta 4; connexin 30.3 [Mus musculus] (SEQ ID NO:63)
- 5) <u>gi|10835079|ref|NP 005259.1|</u> <u>NP 05259</u> gap junction protein, beta 5 (connexin 31.1) [Homo sapiens] (SEQ ID NO:64)
- 6) $\frac{gi|4009522|gb|AAC95472.1|}{AAC95472}$ [AF099731] connexin 31.1 [Homo sapiens] (SEQ ID NO:65)





At page 86 please replace Table 7E with the following:

Applicants: U.S.S.N.:

Guo 09/981,151

Table 7E. Domain Analysis of NOV7

gnl|Pfam|pfam00029, connexin, Connexin. (SEQ ID NO:91)
CD-Length = 218 residues, 100.0% aligned
Score = 318 bits (814), Expect = 3e-88

| Query: | 1 | MNWAFLQGLLSGVNKYSTVLSRIWLSVVFIFRVLVYVVAAEEVWDDEQKDFVCNTKQPGC | 60 |
|------------------|------------|--|------------|
| Sbjct: | 1 | MDWSFLGRLLEGVNKHSTAIGKIWLSVLFIFRILVLGVAAESVWGDEQSDFVCNTQQPGC | 60 |
| Query: | 61 | PNVCYDEFFPVSHVRLWALQLILVTCPSLLVVMHVAYREERERKHHLKHGPNAPSLYDNL | 120 |
| Sbjct: | 61 | + + + + | 120 |
| | | | |
| Query: | 121 | SKKRGGLWWTYLLSLIFKAAVDAGFLYIFHRLYKDYDMPRVVACSVEPC | 169 |
| Query: Sbjct: | 121 121 | SKKRGGLWWTYLLSLIFKAAVDAGFLYIFHRLYKDYDMPRVVACSVEPC + + + + + + + + | 169 179 |
| ~ • | | + | 179 |

At page 87, please replace Table 7F with the following:

Table 7F. Domain Analysis of NOV7

gnl|Smart|smart00037, CNX, Connexin homologues; Connexin channels
participate in the regulation of signaling between developing and
differentiated cell types. (SEQ ID NO:160)
CD-Length = 34 residues, 100.0% aligned
Score = 79.0 bits (193), Expect = 3e-16

At page 100, please replace Table 9C with the following:

| | Table 9C. BLA | AST result | ts for NOV9 | | |
|--|---|----------------|------------------|------------------|--------|
| Gene Index/ Identifier | Protein/ Organism | Length (aa) | Identity (%) | Positives (%) | Expect |
| gi 13591536 emb CAC 36351.1 CAC36351 (AL109613) | dJ1033H22.1 (KIAA0554 protein) [Homo sapiens] | 434 | 373/430 (86%) | 375/430 (86%) | 0.0 |

Applicants: U.S.S.N.:

Guo 09/981,151

| gi 8923249 ref NP 0 60207.1 NP 060207 | hypothetical protein FLJ20275 [Homo sapiens] | 330 | 328/330 (99%) | 329/330 (99%) | e-175 |
|---|--|-----|------------------|------------------|-------|
| gi 10435680 dbj BAB 14638.1 BAB14638 (AK023681) | unnamed protein product [Homo sapiens] | 592 | 319/595 (53%) | 432/595 (71%) | e-160 |
| gi 13936547 gb AAK4 9824.1 AF265550 1 AAK49824 (AF265550) | formin-binding protein 17 [Homo sapiens] | 679 | 307/624 (49%) | 422/624 (67%) | e-148 |
| gi 3043632 dbj BAA2 5480.1 BAA25480 (AB011126) | KIAA0554 protein [Homo sapiens] | 674 | 307/624 (49%) | 422/624 (67%) | e-148 |

At page 100 please replace Table 9D with the following:

Table 9D. ClustalW Analysis of NOV9

| 3) gi 135915 (KIAA0554 pro 4) gi 892324 [Homo sapiens 5) gi 104356 product [Homo 6) gi 139365 binding prote 7) gi 304363 | 9 (SEQ ID NO:24) 36 emb CAC36351 CAC36351 gi 13591536 (AL109613) dJ1033H22.1 tein) [Homo sapiens] (SEQ ID NO:66) 9 ref NP 060207.1 NP 060207 gi 8923249 hypothetical protein FLJ20275] (SEQ ID NO:67) 80 db BAB14638 BAB14638 gi 10435680 (AK023681) unnamed protein sapiens] (SEQ ID NO:68) 47 gb AAK49824.1 AF265550 AAK49824 gi 13936547 (AF265550) forminin 17 [Homo sapiens] (SEQ ID NO:69) 2 db BAA25480.1 BAA25480 gi 3043632 (AB011126) KIAA0554 protein |
|--|--|
| NOV9 gi 13591536 gi 8923249 gi 10435680 gi 13936547 gi 3043632 | 10 20 30 40 LWNGGEEEPPRRPRARSCEPEEAARTPGFPPSRGSRGAKGEEEPPRRPRARSCEPEEAARTPGFPPSRGSRGAKG |
| NOV9 gi 13591536 gi 8923249 gi 10435680 gi 13936547 gi 3043632 | 50 60 70 80 |
| NOV9 g1 13591536 g1 8923249 g1 10435680 g1 13936547 g1 3043632 | 90 100 110 120 |

Applicants: Guo U.S.S.N.: 09/981,151 130 140 160 . | | | | . . NOV9 gi|13591536| gi | 8923249 | SKEEERY OTSCKAFISNLNE NDYAGOHEVOSE SKEEERY OTSCKAFISNLNE NDYAGOHEVOSE SKEEERY OTSCKAFISNLNE NDYAGOHEVOSE gi|10435680| gi | 13936547 | gi|3043632| 200 170 | | | | | | NOV9 YGELMRYAHDLKTERKMHLQEGRKAQQYAHDMCWKQMGNSK gi | 13591536 | gi | 8923249 | IVDLARYVOSLKOERKSNEHDGRKAQOSBETCWKODES IVDLARYVOSLKOERKSNEHDGRKAQOSBETCWKODES IVDLARYVOSLKOERKSNEHDGRKAQOSBETCWKODES gi | 10435680 | gi | 13936547 | gi 3043632 220 230 240 NOV9 gi | 13591536 | gi|8923249| ZEFERDCKEADRAOOYFEKADADINVTKADVEKAROOAOII ZEFERDCKEADRAOOYFEKADADINVTKADVEKAROOAOII ZEFERDCKEADRAOOYFEKADADINVTKADVEKAROOAOII gi|10435680| gi | 13936547 | gi|3043632| 270 280 NOV9 gi | 13591536 | gi | 8923249 | RHOMAGDSKADYSSILOKENBEOHGAAFTHIPNIGOKTOE RHOMAGDSKADYSSILOKENBEOHGAAFTHIPNIGOKTOE RHOMAGDSKADYSSILOKENBEOHGAAFTHIPNIGOKTOE gi 10435680 gi | 13936547 | gi|3043632| 290 320 NOV9 gi | 13591536 | gi | 8923249 | TKLSECYR VRMGESMKTYAE gi | 10435680 | gi | 13936547 | gi 3043632 340 350 360 NOV9 gi | 13591536 | gi 8923249 gi | 10435680 | gi | 13936547 | gi 3043632 370 380 400 390 NOV9 gi|13591536| gi | 8923249 | KGKLW KFG-GKSKGKLWPFIKKN------KFG-GKSKGKLWPFIKKNKLMSLLTS KFG-GKSKGKLWPFIKKNKLMSLLTS gi | 10435680 | gi | 13936547 | GKPDLKFGgi|3043632|

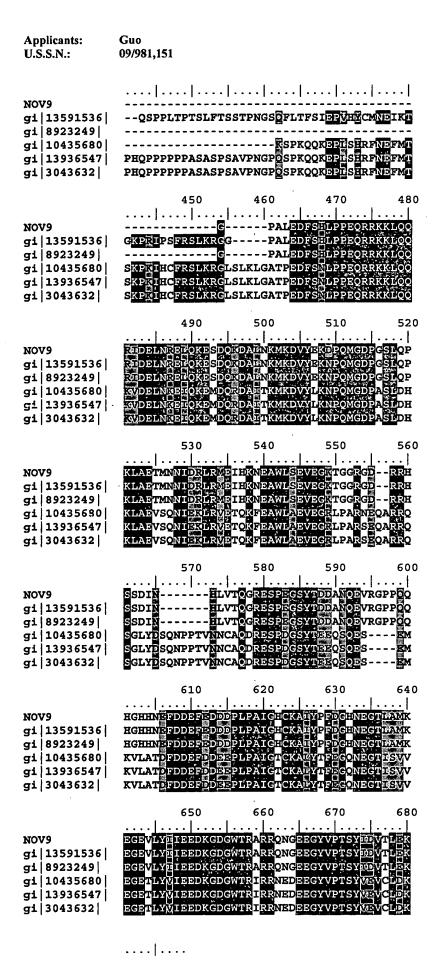
430

440

420

410

- SELVICIONE LA PARTICIONIS.



Applicants: Guo U.S.S.N.: 09/981,151

 NOV9
 NSKGS
 -- (SEQ ID NO:24)

 gi | 13591536 |
 NSKGZVTYI
 (SEQ ID NO:66)

 gi | 8923249 |
 NSKGS
 (SEQ ID NO:67)

 gi | 10435680 |
 NAKCZKTYI
 (SEQ ID NO:68)

 gi | 13936547 |
 NAKOS
 (SEQ ID NO:69)

 gi | 3043632 |
 NAKOS
 (SEQ ID NO:70)

At page 103, please replace Table 9E with the following:

Table 9E. Domain Analysis of NOV9

gnl|Smart|smart00326,-SH3, Src homology 3 domains; Src homology 3
(SH3) domains bind to target proteins through sequences containing
proline and hydrophobic amino acids. Pro-containing polypeptides may
bind to SH3 domains in 2 different binding orientations. (SEQ ID
NO:92)

CD-Length = 59 residues, 88.1% aligned Score = 64.7 bits (156), Expect = 1e-11

Query: 484 HCKAIYPFDGHNEGTLAMKEGEVLYIIEEDKGDGWTRARRQNGEEGYVPTSYI 536

Sbjct: 4 QVRALYDYTAQDPDELSFKKGDIITVLEKS-DDGWWKGRLGTGKEGLFPSNYV 55

Query: (of SEQ ID NO:24)

Sbjct: (SEQ ID NO:92)

At page 104, please replace Table 9F with the following:

Table 9F. Domain Analysis of NOV9

gnl|Pfam|pfam00018, SH3, SH3 domain. SH3 (Src homology 3) domains are
often indicative of a protein involved in signal transduction related
to cytoskeletal organization. First described in the Src cytoplasmic
tyrosine kinase. The structure is a partly opened beta barrel. (SEQ
ID NO:93)

CD-Length = 57 residues, 91.2% aligned Score = 63.5 bits (153), Expect = 3e-11

Query: 486 KAIYPFDGHNEGTLAMKEGEVLYIIEEDKGDGWTRARRQNGEEGYVPTSYID 53

A+Y + L+ K+G+++ ++E+ GW + R + +EG +P++Y++

Sbjct: 4 VALYDYQARESDELSFKKGDIIIVLEKSDDGGWWKGRLKGTKEGLIPSNYVE 55

Query: of SEQ ID NO:24)

Sbjct: (SEQ ID NO:93)

Applicants: Guo U.S.S.N.: 09/981,151

At page 104, please replace Table 9 with the following:

Table 9G. Domain Analysis of NOV9

gnl|Smart|smart00055, FCH, Fes/CIP4 homology domain; Alignment
extended from original report. Highly alpha-helical. Also known as the
RAEYL motif or the S. pombe Cdc15 N-terminal domain. (SEQ ID NO:94)
CD-Length = 91 residues, 97.8% aligned
Score = 58.2 bits (139), Expect = 1e-09

At page 104, please replace Table 9H with the following:

Table 9H. Domain Analysis of NOV9

gnl|Pfam|pfam00611, FCH, Fes/CIP4 homology domain. Alignment extended from. Highly alpha-helical. (SEQ ID NO:95) CD-Length = 94 residues, 97.9% aligned Score = 40.0 bits (92), Expect = 3e-04

At page 113 please replace Table 10F with the following:

Applicants:

Guo 09/981,151

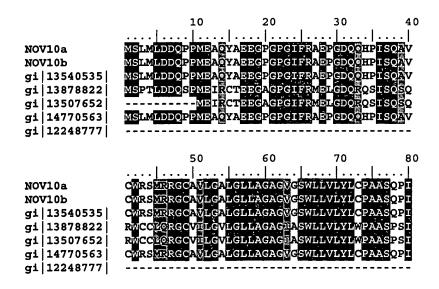
U.S.S.N.:

| | Table 10F. BLA | ST result | s for NOV10 | a | |
|--|--|----------------|------------------|-------------------|--------|
| Gene Index/ Identifier | Protein/ Organism | Length (aa) | Identity (%) | Positives (%) | Expect |
| <u>gi 13540535 ref NP</u> <u>110397.1 </u> NP 110397 | transmembrane protease, serine 5; spinesin [Homo sapiens] | 457 | 452/463 (97%) | 453/463 (97%) | 0.0 |
| gi 13878822 sp Q9ER 04 TM65-MOUSE Q9ER04 | TRANSMEMBRANE PROTEASE, SERINE 5 (SPINESIN) | 455 | 360/461 (78%) | 394/461 (85%) | 0.0 |
| gi 13507652 ref NP 109634.1 NP 109634 | transmembrane protease, serine 5 (spinesin) [Mus musculus] | 445 | 354/451 (78%) | 388/451 (85%), | 0.0 |
| gi 14770563 ref XP 041427.1 -XP 041427 | transmembrane protease, serine 5 [Homo sapiens] | 398 | 354/362 (97%) | 355/362 (97%) | 0.0 |
| gi 12248777 dbj BAB 20276.1 BAB20276 (AB016229) | type 2 spinesin [Mus musculus] | 311 | 260/317 (82%) | 281/317 (88%) | e-146 |

At page 114 please replace Table 10G with the following:

Table 10G. ClustalW Analysis of NOV10

- Novel NOV10a (SEQ ID NO:27) 1)
- (SEQ ID NO:29) Novel NOV10b 2)
- gi | 13540535 | ref | NP 110397.1 | NP 110397 gi | 13540535 | transmembrane protease, serine 5; spinesin [Homo sapiens] (SEQ ID NO:71)
- 5) qi|13878822|sp|Q9ER04|TMS5-MOUSE Q9ER04 **gi|13878822**| TRANSMEMBRANE PROTEASE, SERINE 5 (SPINESIN) (SEQ ID NO:72)
- 6) gi|13507652|ref|NP-109634.1| NP 109634 gi|13507652| transmembrane protease, serine 5 (spinesin) [Mus musculus] (SEQ ID NO:73)
- gi | 14770563 | ref | XP 041427.1 | XP 041427 gi | 14770563 | transmembrane protease, serine 5 [Homo sapiens] (SEQ ID NO:74)
- 8) gi|12248777|dbj|BAB20276.1| BAB20276 gi|12248777| (AB016229) type 2 spinesin [Mus musculus] (SEQ ID NO:75)



Applicants: Guo U.S.S.N.: 09/981,151 120 90 NOV10a NOV10b gi | 13540535 | gi | 13878822 | gi | 13507652 | gi | 14770563 | gi | 12248777 | 160 NOV10a NOV10b gi|13540535| gi | 13878822 | gi | 13507652 | gi | 14770563 | gi | 12248777 | NOV10a NOV10b gi|13540535| AVNLSDÍKLMRSOEFAÖLSARPGGLVEEAW AVNLSDÍKLMRSOEFAÖLSARPGGLVEEAW GVNLSDÍKLMSSOEFAÖLSPRIGGEFEEAW AVNLSDÍKLMRSOEFAOLSARPGGLVEEAW gi|13878822| gi | 13507652 | gi|14770563| gi | 12248777 | 240 NOV10a NOV10b gi | 13540535 | gi | 13878822 | gi | 13507652 | CSECGARPLASRIVGO gi | 14770563 | SGR VSL CSECGARPLASRIVGGO VASGRWPWOA gi | 12248777 |

NOV10a NOV10b gi|13540535| gi | 13878822 | gi | 13507652 | gi | 14770563 |

NOV10a

NOV10b

NOV10a

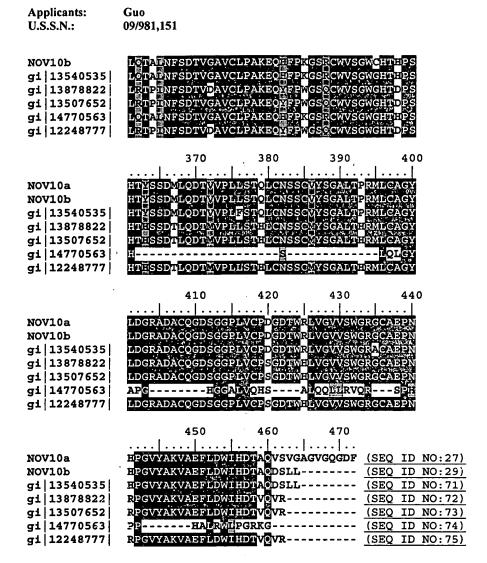
gi | 12248777 |



LGSRHTCGASVLAPEWVVTAAHCMASFRL

290

320







At page 116, please replace Table 10H with the following:

Table 10H Domain Analysis of NOV10

gnl|Smart|smart00020, Tryp_SPc, Trypsin-like serine protease; Many of
these are synthesised as inactive precursor zymogens that are cleaved
during limited proteolysis to generate their active forms. A few,
however, are active as single chain molecules, and others are inactive
due to substitutions of the catalytic triad residues. (SEQ ID NO:96)
CD-Length = 230 residues, 100.0% aligned
Score = 266 bits (681), Expect = 2e-72

| | 224 1 | RIVGGQSVAPGRWPWQASVAL-GFRHTCGGSVLAPRWVVTAAHCMHSFRLARLSSWRVHA | 282 57 |
|--------|----------|--|------------|
| - • | | GLVSHSAVRPHQGALVERIIPHPLYSAQNHDYDVALLRLQTALNFSDTVGAVCLPAKEQH | 342 117 |
| Query: | 343 | FPKGSRCWVSGWGHTHPSHTYSSDMLQDTVVPLLSTQLCNSSCVYSGALTPRMLCAGYLD | 402 |

| U.S.S.N.: | | • | 09/981,151 | | | | |
|-----------|--------|------|--|---|--|--|--|
| | Sbjct: | 118 | + + + + + | 7 | | | |
| | Query: | 403 | GRADACQGDSGGPLVCPDGDTWRLVGVVSWGR-GCAEPNHPGVYAKVAEFLDWI 455 | | | | |
| | Sbjct: | 178 | + + + + + GGKDACQGDSGGPLVC-NDPRWVLVGIVSWGSYGCARPNKPGVYTRVSSYLDWI 230 | | | | |
| | Query: | (of | SEQ ID NO:27) | | | | |
| | Sbjct: | (SEQ | ID NO:96) | | | | |

At page 116, please replace Table 10I with the following:

Table 10I. Domain Analysis of NOV10

gnl | Pfam | pfam00089, trypsin, Trypsin. Proteins recognized include all proteins in families S1, S2A, S2B, S2C, and S5 in the classification of peptidases. Also included are proteins that are clearly members, but that lack peptidase activity, such as haptoglobin and protein Z (PRTZ*). (SEQ ID NO:97)

CD-Length = 217 residues, 100.0% aligned core = 211 bits (538), Expect = 6e-56

| Query: | 225 | <pre>IVGGQSVAPGRWPWQASVALGFRHTCGGSVLAPRWVVTAAHCMHSFRLARLSSWRVHAGL +</pre> | 284 |
|----------|-----|---|-----|
| Sbjct: | 1 | IVGGREAQAGSFPWQVSLQVSSGHFCGGSLISENWVLTAAHCVSGASSVRVVLGE | 55 |
| Query: . | 285 | VSHSAVRPH-QGALVERIIPHPLYSAQNHDYDVALLRLQTALNFSDTVGAVCLPAKEQHF + | 343 |
| Sbjct: | 56 | HNLGTTEGTEQKFDVKKIIVHPNYNPDTNDIALLKLKSPVTLGDTVRPICLPSASSDL | 113 |
| Query: | 344 | PKGSRCWVSGWGHTHPSHTYSSDMLQDTVVPLLSTQLCNSSCVYSGALTPRMLCAGYLDG | 403 |
| Sbjct: | 114 | | 169 |
| Query: | 404 | RADACQGDSGGPLVCPDGDTWRLVGVVSWGRGCAEPNHPGVYAKVAEFLDWI 455 | |
| Sbjct: | 170 | K-DACQGDSGGPLVCSDGELVGIVSWGYGCAVGNYPGVYTRVSRYLDWI 217 | |
| Query: | (of | SEQ ID NO:27) | |
| | | | |

Sbjct: (SEQ ID NO:97)

Applicants:

Guo